EAST Search History

| Ref # | Hits | Search Query | DBs | Default Operator | Plurals | Time Stamp |
|----------|---------|---|--|---------------------|---------|------------------|
| L1 | 1433 | 514/649 | US-PGPUB; USPAT; EPO; DERWENT | OR | ON | 2006/11/02 16:59 |
| L2 | 3277 | 424/486 | US-PGPUB; USPAT; EPO; DERWENT | OR | ON | 2006/11/02 16:59 |
| L3 | 14 | L1 and L2 | US-PGPUB; USPAT; EPO; DERWENT | OR | ON | 2006/11/02 16:59 |
| L4 | 27 | Fesoterodine . | US-PGPUB; USPAT; EPO; DERWENT | OR | ON | 2006/11/02 17:01 |
| L5 | 2283 | 514/249 | US-PGPUB; USPAT; EPO; DERWENT | OR | ON | 2006/11/02 17:01 |
| L6 | 2 | L4 and L5 | US-PGPUB; USPAT; EPO; DERWENT | OR | ON | 2006/11/02 17:01 |
| S1 | 5547820 | (R)-2-[3-(1, 1-diisopropylamino)-1-phenylpropyl]-4-(hyd roxymethyl)phenyl isobutyrate | US-PGPUB; USPAT; EPO; DERWENT | OR | ON | 2006/11/02 11:51 |
| S2 | 27 | Fesoterodine | US-PGPUB; USPAT; EPO; DERWENT | OR | ON | 2006/11/02 11:51 |
| S3 | . 27 | S1 and S2 | US-PGPUB; USPAT; EPO; DERWENT | OR | ON | 2006/11/02 16:59 |

11/2/2006 5:02:37 PM Page 1

Welcome to STN International! Enter x:x

LOGINID:ssptalxn1621

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

Welcome to STN International

```
NEWS 1
                Web Page URLs for STN Seminar Schedule - N. America
NEWS 2
                "Ask CAS" for self-help around the clock
                INSPEC enhanced with 1898-1968 archive
NEWS 3 AUG 09
NEWS 4 AUG 28 ADISCTI Reloaded and Enhanced
NEWS 5 AUG 30 CA(SM)/CAplus(SM) Austrian patent law changes
NEWS 6 SEP 11 CA/Caplus enhanced with more pre-1907 records
NEWS 7 SEP 21 CA/CAplus fields enhanced with simultaneous left and right
                truncation
                CA(SM)/CAplus(SM) display of CA Lexicon enhanced
NEWS 8 SEP 25
NEWS 9 SEP 25
               CAS REGISTRY(SM) no longer includes Concord 3D coordinates
NEWS 10 SEP 25
               CAS REGISTRY(SM) updated with amino acid codes for pyrrolysine
NEWS 11
       SEP 28
               CEABA-VTB classification code fields reloaded with new
                classification scheme
NEWS 12 OCT 19
                LOGOFF HOLD duration extended to 120 minutes
NEWS 13 OCT 19
                E-mail format enhanced
                Option to turn off MARPAT highlighting enhancements available
NEWS 14 OCT 23
NEWS 15 OCT 23
                CAS Registry Number crossover limit increased to 300,000 in
                multiple databases
NEWS 16
        OCT 23
                The Derwent World Patents Index suite of databases on STN
```

has been enhanced and reloaded

OCT 30 CHEMLIST enhanced with new search and display field NEWS 17

JUNE 30 CURRENT WINDOWS VERSION IS V8.01b, CURRENT NEWS EXPRESS MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP), AND CURRENT DISCOVER FILE IS DATED 26 JUNE 2006.

STN Operating Hours Plus Help Desk Availability NEWS HOURS Welcome Banner and News Items NEWS LOGIN For general information regarding STN implementation of IPC 8 NEWS IPC8 NEWS X25 X.25 communication option no longer available

Enter NEWS followed by the item number or name to see news on that specific topic.

All use of STN is subject to the provisions of the STN Customer agreement. Please note that this agreement limits use to scientific research. Use for software development or design or implementation of commercial gateways or other similar uses is prohibited and may result in loss of user privileges and other penalties.

* * * * * * * * * * STN Columbus * * * * * * *

FILE 'HOME' ENTERED AT 12:22:33 ON 02 NOV 2006

=> file caplus COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.21 0.21

FULL ESTIMATED COST

FILE 'CAPLUS' ENTERED AT 12:22:47 ON 02 NOV 2006 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2006 AMERICAN CHEMICAL SOCIETY (ACS)

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the American Chemical Society and is provided to assist you in searching databases on STN. Any dissemination, distribution, copying, or storing of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 2 Nov 2006 VOL 145 ISS 19 FILE LAST UPDATED: 1 Nov 2006 (20061101/ED)

Effective October 17, 2005, revised CAS Information Use Policies apply. They are available for your review at:

http://www.cas.org/infopolicy.html

=>

Uploading C:\Program Files\Stnexp\Queries\fesoterodine.str

chain nodes :

7 8 9 10 11 12 13 14 15 16 23 24 25 26 27 28 29

ring nodes :

1 2 3 4 5 6 17 18 19 20 21 22

chain bonds :

5-7 7-8 7-17 8-9 9-10 10-11 10-12 11-15 11-16 12-13 12-14 19-27 22-23

23-24 24-25 24-26 27-28 28-29

ring bonds :

1-2 1-6 2-3 3-4 4-5 5-6 17-18 17-22 18-19 19-20 20-21 21-22

exact/norm bonds :

9-10 10-11 10-12 22-23 23-24 24-26 27-28

exact bonds :

5-7 7-8 7-17 8-9 11-15 11-16 12-13 12-14 19-27 24-25 28-29

normalized bonds :

1-2 1-6 2-3 3-4 4-5 5-6 17-18 17-22 18-19 19-20 20-21 21-22

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:CLASS 8:CLASS 9:CLASS 10:CLASS 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:CLASS 17:Atom 18:Atom 19:Atom 20:Atom 21:Atom 22:Atom 23:CLASS 24:CLASS 25:CLASS 26:CLASS 27:CLASS 28:CLASS 29:CLASS

STRUCTURE UPLOADED L1

=> d L1

L1 HAS NO ANSWERS

L1STR

Structure attributes must be viewed using STN Express query preparation.

=> s L1

REG1stRY INITIATED

Substance data SEARCH and crossover from CAS REGISTRY in progress... Use DISPLAY HITSTR (or FHITSTR) to directly view retrieved structures.

SAMPLE SEARCH INITIATED 12:23:18 FILE 'REGISTRY' SAMPLE SCREEN SEARCH COMPLETED -5 TO ITERATE

100.0% PROCESSED

5 ITERATIONS

0 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH **COMPLETE**

PROJECTED ITERATIONS:

5 TO 234

PROJECTED ANSWERS:

0 TO

L2 0 SEA SSS SAM L1

L3 0 L2

=> file caplus

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION

1.57 FULL ESTIMATED COST 0.46

FILE 'CAPLUS' ENTERED AT 12:23:40 ON 02 NOV 2006 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2006 AMERICAN CHEMICAL SOCIETY (ACS)

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the American Chemical Society and is provided to assist you in searching databases on STN. Any dissemination, distribution, copying, or storing of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 2 Nov 2006 VOL 145 ISS 19 FILE LAST UPDATED: 1 Nov 2006 (20061101/ED)

Effective October 17, 2005, revised CAS Information Use Policies apply. They are available for your review at:

http://www.cas.org/infopolicy.html

=> s L2

L4 0 L2

=> s fesoterodine

L5 9 FESOTERODINE

=> d L5 1-9 fhit ibib abs
'FHIT' IS NOT A VALID FORMAT FOR FILE 'CAPLUS'

The following are valid formats:

ABS ----- GI and AB

ALL ----- BIB, AB, IND, RE

APPS ----- AI, PRAI

BIB ----- AN, plus Bibliographic Data and PI table (default) CAN ----- List of CA abstract numbers without answer numbers

CBIB ----- AN, plus Compressed Bibliographic Data

CLASS ----- IPC, NCL, ECLA, FTERM

DALL ----- ALL, delimited (end of each field identified)

DMAX ----- MAX, delimited for post-processing

FAM ----- AN, PI and PRAI in table, plus Patent Family data

FBIB ----- AN, BIB, plus Patent FAM

IND ----- Indexing data

IPC ------ International Patent Classifications

MAX ----- ALL, plus Patent FAM, RE

PATS ----- PI, SO

SAM ----- CC, SX, TI, ST, IT

SCAN ----- CC, SX, TI, ST, IT (random display, no answer numbers; SCAN must be entered on the same line as the DISPLAY,

e.g., D SCAN or DISPLAY SCAN)

STD ----- BIB, CLASS

IABS ----- ABS, indented with text labels

IALL ----- ALL, indented with text labels

IBIB ----- BIB, indented with text labels

IMAX ----- MAX, indented with text labels
ISTD ----- STD, indented with text labels

OBIB ----- AN, plus Bibliographic Data (original)

OIBIB ----- OBIB, indented with text labels

SBIB ----- BIB, no citations SIBIB ----- IBIB, no citations

HIT ----- Fields containing hit terms

HITIND ----- IC, ICA, ICI, NCL, CC and index field (ST and IT) containing hit terms

HITRN ----- HIT RN and its text modification

HITSTR ----- HIT RN, its text modification, its CA index name, and its structure diagram

HITSEQ ----- HIT RN, its text modification, its CA index name, its structure diagram, plus NTE and SEQ fields

FHITSTR ---- First HIT RN, its text modification, its CA index name, and its structure diagram

FHITSEQ ---- First HIT RN, its text modification, its CA index name, its structure diagram, plus NTE and SEQ fields

KWIC ----- Hit term plus 20 words on either side

OCC ----- Number of occurrence of hit term and field in which it occurs

To display a particular field or fields, enter the display field codes. For a list of the display field codes, enter HELP DFIELDS at an arrow prompt (=>). Examples of formats include: TI; TI,AU; BIB,ST; TI,IND; TI,SO. You may specify the format fields in any order and the information will be displayed in the same order as the format specification.

All of the formats (except for SAM, SCAN, HIT, HITIND, HITRN, HITSTR, FHITSTR, HITSEQ, FHITSEQ, KWIC, and OCC) may be used with DISPLAY ACC to view a specified Accession Number.
ENTER DISPLAY FORMAT (BIB):bib abs

- L5 ANSWER 1 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
- AN 2006:1133705 CAPLUS
- TI Treatment of the overactive bladder syndrome with muscarinic receptor antagonists a matter of metabolites?
- AU Michel, Martin C.; Hegde, Sharath S.
- CS Department of Pharmacology & Pharmacotherapy, Academic Medical Center, University of Amsterdam, Meibergdreef 15, Amsterdam, 1105 AZ, Neth.
- SO Naunyn-Schmiedeberg's Archives of Pharmacology (2006), 374(2), 79-85 CODEN: NSAPCC; ISSN: 0028-1298
- PB Springer
- DT Journal
- LA English
- Antagonists of muscarinic acetylcholine receptors, such as darifenacin, AΒ oxybutynin, propiverine, solifenacin, tolterodine, and trospium, are the mainstay of the treatment of the overactive bladder syndrome. Fesoterodine is a newer drug awaiting regulatory approval. We briefly review the pharmacol. activity of their metabolites and discuss how active metabolites may contribute to their efficacy and tolerability in vivo. Except for trospium, and perhaps solifenacin, all of the above drugs form active metabolites, and their presence and activity need to be taken into consideration when elucidating relationships between pharmacokinetics and pharmacodynamics of these drugs. Moreover, the ratios between parent compds. and metabolites may differ depending on genotype of the metabolizing enzymes, concomitant medication, and/or drug formulation. Differential generation of active metabolites of darifenacin or tolterodine are unlikely to influence the overall clin. profile of these drugs in a major way because the active metabolites exhibit a similar pharmacol. profile as the parent compound In contrast, metabolites of oxybutynin and propiverine may behave quant. or even qual. differently from their parent compds. and this may have an impact on the overall clin. profile of these drugs. We conclude that more comprehensive studies of drug metabolites are required for an improved understanding of their clin. effects.

```
DN
     144:156740
     Combinations of statins with bronchodilators for treatment of respiratory
TI
     disorders
     Lindmark, Bertil; Thoren, Anders Ingemar
IN
     AstraZeneca AB, Swed.; AstraZeneca UK Limited
PΑ
     PCT Int. Appl., 18 pp.
SO
     CODEN: PIXXD2
     Patent
DT
     English
LA
FAN.CNT 1
                                            APPLICATION NO.
                                                                   DATE
     PATENT NO.
                         KIND
                                DATE
                                                                   --------
                                -----
                                            -----
     _____
                         ____
                                            WO 2005-GB2413 .
                                                                    20050620
                                20060126
     WO 2006008437
ΡI
                         A1
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
             CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
             GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ,
             LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA,
             NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK,
             SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU,
             ZA, ZM, ZW
         RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE,
             IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF,
             CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM,
             KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG,
             KZ, MD, RU, TJ, TM
                                20040715
PRAI GB 2004-15789
                          Α
     The invention provides medicaments comprising combinations of
     bronchodilators, glucocorticosteroids and HMG-CoA reductase inhibitors in
     the treatment of respiratory disorders such as chronic obstructive
    pulmonary disease (COPD). For example, a metered dose inhaler contained
    per dose formoterol fumarate dihydrate 4.5 μg, budesonide 160 μg,
     rosuvastatin 1 mg, and HFA 227 50 µL. Also, an inhalation/oral
     combination comprised an aerosol formulation containing per dose formoterol
     fumarate dihydrate 4.5 \mu g and budesonide 160 \mu g, and a tablet
     formulation containing rosuvastatin 10 mg.
              THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT 3
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
     ANSWER 3 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
L5
ΑN
     2004:878361 CAPLUS
DN
     141:370546
     Highly pure bases of 3,3-diphenyl propylamine monoesters for use in
ΤI
     transdermal delivery systems
     Breitenbach, Armin; Meese, Claus; Wolff, Hans-Michael; Drews, Roland
IN
     Schwarz Pharma Ag, Germany
PA
SO
     PCT Int. Appl., 72 pp.
     CODEN: PIXXD2
DT
     Patent
     German
LΑ
FAN.CNT 1
                                                                   DATE
                                DATE .
                                           APPLICATION NO.
     PATENT NO.
                         KIND
                                            ______
                                -----
     _____
                         ----
                                          WO 2004-EP3567
                                                                    20040403
                                20041021
PΙ
     WO 2004089872
                         A1
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
             CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
             GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
             LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
             NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
         TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
             BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,
             ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI,
             SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN,
             TD, TG
```

2006:76147 CAPLUS

AN

| | DE | 1031 | 5917 | | | A1 | - 2 | 2004 | 1118 | DE | 20 | 003- | 1031 | 5917 | | 2 | 0030 | 408 | |
|------|-------------------|------|-------|------|------|------|--------------------|-----------------------|------|-------|----------|----------|----------|------|-----|-----|------|-----|----|
| | AU 2004228163 | | | A1 | 2 | 2004 | 1021 | 21 AU 2004-228163 | | | | 63 | 20040403 | | | | | | |
| | CA 2505848 | | | AA | - 2 | 2004 | 1021 | CA 2004-2505848 | | | | 20040403 | | | | | | | |
| | BR 2004006221 | | | Α | 2 | 2005 | 0809 | BR 2004-6221 | | | | 20040403 | | | | | | | |
| | EP 1613584 | | A1 | 2 | 2006 | 0111 | 111 EP 2004-725610 | | | | 20040403 | | | | | | | | |
| | | R: | AT, | BE, | CH, | DE, | DK, | ES, | FR, | GB, G | R, | IT, | LI, | LU, | NL, | SE, | MC, | PT, | |
| | | | ΙE, | SI, | LT, | LV, | FI, | RO, | MK, | CY, A | L, | TR, | BG, | CZ, | EE, | HU, | PL, | SK, | HR |
| | CN 1802345 | | Α | 2 | 2006 | 0712 | CN | CN 2004-80009224 2004 | | | | 0040 | 403 | | | | | | |
| | JΡ | 2006 | 5227 | 58 | | T2 | 2 | 2006 | 1005 | JP | 20 | 006- | 5049 | 89 | | 2 | 0040 | 403 | |
| | US | 2006 | 01483 | 32 | • | A1 | 2 | 2006 | 0119 | US | 20 | 005- | 5328 | 36 | | 2 | 0050 | 426 | |
| | NO | 2005 | 0050 | 78 | | Α | 2 | 2005 | 1031 | NO | 20 | 005- | 5078 | | | 2 | 0051 | 031 | |
| PRAI | DΕ | 2003 | -103 | 1591 | 7 | Α | 2 | 2003 | 0408 | | | | | | | | | | |
| | WO | 2004 | -EP3 | 567 | | W | 2 | 2004 | 0403 | | | | | | | | | | |
| os | MARPAT 141:370546 | | | | | | | | | | | | | | | | | | |
| GT | | | | | | | | | | | | | | | | | | | |

The invention relates to a compound of general formula (I) wherein A AΒ represents deuterium or hydrogen, R represents a group selected from C1-6 alkyl, C3-10 cycloalkyl or Ph, which can be substituted by C1-3 alkoxy, fluorine, chlorine, bromine, iodine, nitro, amino, hydroxyl, oxo, mercapto or deuterium. The C atom marked with a * (star) can be present in an (R) configuration, in an (S)-configuration or a mixture thereof. The invention is characterized in that the above-mentioned compds. are free bases with a degree of purity of more than 97 wt %. The invention also relates to a method for the production of highly pure compds. of general formula (I) and to the use thereof in the production of medicaments. Thus (R)-2-[3-(Diisopropylamino) - 1 - phenylpropyl] - 4 - (hydroxymethyl) phenol was reacted with isobutyric acid chloride to form fesoterodine. Fesoterodine was purified via the formation of its fumaric acid 1.5 G of the highly pure fesoterodine was mixed with 8.5 g silicone adhesive Bio-PSA 7-4300 and applied to a foil in order to prepare a transdermal delivery system.

RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

```
L5 ANSWER 4 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
```

AN 2004:875349 CAPLUS

DN 142:303234

TI Mucosal adjuvants and delivery systems for oral and nasal vaccination

AU Baudner, Barbara C.; Verhoel, J. Coos; Junginger, Hans E.; del Giudice, Giuseppe

CS IRIS Research Center, Siena, 53100, Italy

SO Drugs of the Future (2004), 29(7), 721-732 CODEN: DRFUD4; ISSN: 0377-8282

PB Prous Science

DT Journal; General Review

LA English

A review. The pillars of pharmacotherapy for overactive bladder (OAB) are AB antimuscarinic agents which inhibit bladder smooth muscle contractions through interference with acetylcholine action on muscarinic receptors of the detrusor smooth muscle. Despite the availability of different antimuscarinic compds., physicians and patients remain dissatisfied with current treatments due to adverse events and/or insufficient efficacy. Therefore, new agents with improved safety and efficacy profiles are needed for a more effective treatment of overactive bladder. Fesoterodine is a novel bladder-selective muscarinic antagonist that has shown potent antimuscarinic activity in vitro and in vivo. In multiple investigations, the agent has been shown to be safe and well tolerated in subjects of different ethnic origin, age and gender; in poor and extensive CYP2D6 metabolizers; in subjects taking concomitant medication inhibiting CYP3A4; in fed or fasted states; and in those suffering from hepatic impairment. No clin. relevant changes in heart rate, blood pressure, ECG parameters or laboratory analyses have been seen with therapeutic doses of fesoterodine in these studies. Furthermore, in a phase II clin. trial in patients with OAB, fesoterodine demonstrated rapid and significant efficacy on a variety of endpoints. The results of this trial encouraged the manufacturer (SCHWARZ PHARMA) to initiate a phase III clin. trial program for fesoterodine.

RE.CNT 169 THERE ARE 169 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L5 ANSWER 5 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
- AN 2004:875348 CAPLUS
- DN 142:147630
- TI Fesoterodine, an advanced antimuscarinic for the treatment of overactive bladder: a safety update
- AU Cole, Patrick
- CS Medical Information Dept., Prous Science, Barcelona, 08080, Spain
- SO Drugs of the Future (2004), 29(7), 715-720 CODEN: DRFUD4; ISSN: 0377-8282
- PB Prous Science
- DT Journal; General Review
- LA English
- The pillars of pharmacotherapy for overactive bladder (OAB) are AB antimuscarinic agents which inhibit bladder smooth muscle contractions through interference with acetylcholine action on muscarinic receptors of the detrusor smooth muscle. Despite the availability of different antimuscarinic compds., physicians and patients remain dissatisfied with current treatments due to adverse events and/or insufficient efficacy. Therefore, new agents with improved safety and efficacy profiles are needed for a more effective treatment of overactive bladder. Fesoterodine is a novel bladder-selective muscarinic antagonist that has shown potent antimuscarinic activity in vitro and in vivo. In multiple investigations, the agent has been shown to be safe and well tolerated in subjects of different ethnic origin, age and gender; in poor and extensive CYP2D6 metabolizers; in subjects taking concomitant medication inhibiting CYP3A4; in fed or fasted states; and in those suffering from hepatic impairment. No clin. relevant changes in heart rate, blood pressure, ECG parameters or laboratory analyses have been seen with therapeutic doses of fesoterodine in these studies. Furthermore, in a phase II clin. trial in patients with OAB, fesoterodine demonstrated rapid and significant efficacy on a variety of endpoints. The results of this trial encouraged the manufacturer (SCHWARZ PHARMA) to initiate a phase III clin. trial program for fesoterodine.
- RE.CNT 24 THERE ARE 24 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L5 ANSWER 6 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
- AN 2004:872676 CAPLUS
- DN 141:337790

```
TI
     Transdermal administration of (R)-3,3-diphenylpropylamine monoesters
     Breitenbach, Armin; Meese, Claus; Wolff, Hans-Michael; Drews, Roland
IN
PA
     Schwarz Pharma Ag, Germany
     PCT Int. Appl., 68 pp.
SO
     CODEN: PIXXD2
     Patent
DT
     German
LA
FAN.CNT 1
                                            APPLICATION NO.
                                                                    DATE
                                DATE
     PATENT NO.
                         KIND
                                            -----
     _____
                                -----
                         ----
     WO 2004089346
                                20041021
                                            WO 2004-EP3574
                                                                    20040403
ΡI
                         A1
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
             CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
             GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
             LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
             NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
             TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
        RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
             BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,
             ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI,
             SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN,
             TD, TG
                                                                    20030408
                                20041104
                                            DE 2003-10315878
    DE 10315878
                          Αl
                                            AU 2004-228927
                                                                    20040403
    AU 2004228927
                          A1
                                20041021
                                                                    20040403
                                            CA 2004-2505780
                          AA
                                20041021
     CA 2505780
                                            EP 2004-725614
                                                                    20040403
                                20050518
    EP 1530461
                          A1
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK,
    BR 2004006212
                          Α
                                20050816
                                            BR 2004-6212
                                                                    20040403
                                            CN 2004-80009176
                                                                    20040403
     CN 1767820
                          Α
                                20060503
                                                                    20040403
                                            JP 2006-504992
     JP 2006522759
                          T2
                                20061005
                                                                    20050401
                                            ZA 2005-2681
     ZA 2005002681
                          Α
                                20051013
                                                                    20050426
                                            US 2005-533683
     US 2006029673
                          A1
                                20060209
                                                                    20051010
                                            NO 2005-4644
    NO 2005004644
                          Α
                                20051010
PRAI DE 2003-10315878
                          Α
                                20030408
                                20040403
     WO 2004-EP3574
                          W
OS
     MARPAT 141:337790
```

GI

The invention relates to a device for transdermally administering a compound of formula (I), wherein A represents hydrogen or deuterium, R represents a group selected among C1-6 alkyl, C3-10 cycloalkyl, or Ph, each of which can be substituted by C1-3 alkoxy, fluoride, chlorine, bromine, iodine, nitro, amino, hydroxy, oxo, mercapto, or deuterium, the C atom marked by * (asterisk) being provided in the R configuration. The invention is characterized in that the compound of general formula (I) is provided in a polymer matrix and is released at a dose of 0.5 to 20 mg per day through human skin. The invention further relates to the use of said compds. of

formula (I) for producing transdermal medicaments. Thus a silicone-based transdermal system was prepared by the hot-melt process. 8.5 G of an adhesive mixture composed of BIO-PSA 7-4300 from Dow-Corning and 5

weight/weight%

ozokerite or ceresin was heated to 150°C for 20 min until a homogeneous melt was formed. 1.5 G fesoterodine were added to the melt; the mixture was kept for addnl. 5 min at 150°C; followed by application onto a preheated foil. 5 Cm2 samples were used for dissoln. studies.

- RE.CNT 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L5 ANSWER 7 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
- AN 2004:761399 CAPLUS
- DN 141:254396
- TI Fesoterodine a new effective and well-tolerated antimuscarinic for the treatment of urgency-frequency syndrome: results of a phase 2 controlled study
- CS Chapple C1, Royal Hallamshire Hospital, UK
- SO Neurourology and Urodynamics (2004), 23(5/6), 598-599 CODEN: NEUREM; ISSN: 0733-2467
- PB Wiley-Liss, Inc.
- DT Journal
- LA English
- AB Fesoterodine as new effective and well-tolerated antimuscarinic for the treatment of urgency-frequency syndrome is studied here.
- L5 ANSWER 8 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
- AN 2003:993805 CAPLUS
- DN 140:331551
- TI Fesoterodine: Treatment of urinary incontinence muscarinic M3 antagonist
- AU Sorbera, L. A.; Castaner, J.; Lesson, P. A.
- CS Prous Science, Barcelona, 08080, Spain
- SO Drugs of the Future (2003), 28(7), 647-651 CODEN: DRFUD4; ISSN: 0377-8282
- PB Prous Science
- DT Journal; General Review
- LA English
- Urinary incontinence and overactive bladder are extremely AB A review. common disorders affecting up to 12 and 20 million adults in the U.S., resp. Current pharmacotherapy includes peripherally acting compds. which modulate bladder smooth muscle contraction or centrally acting agents which modulate the neurol. control of urination. Anticholinergic agents inhibit bladder smooth muscle contraction through interference with acetylcholine action on muscarinic receptors on detrusor smooth muscle. However, the first anticholinergic agents were associated with a high rate of adverse events due to nonselectivity and targeting of several muscarinic subtypes and thus other organs. The search for novel, more bladder-selective antimuscarinic agents with better tolerability was initiated. Fesoterodine is a novel selective muscarinic M3 receptor antagonist that has shown potent antimuscarinic activity in vitro and in vivo and has been selected for further development as a treatment for urinary incontinence and overactive bladder.
- RE.CNT 14 THERE ARE 14 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L5 ANSWER 9 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
- AN 2003:950829 CAPLUS
- DN 140:13084
- TI Combination of selected opioids with other active substances for use in the therapy of urinary incontinence
- IN Christoph, Thomas
- PA Grunenthal G.m.b.H., Germany
- SO PCT Int. Appl., 126 pp.

CODEN: PIXXD2 DT Patent German FAN.CNT 1 KIND DATE APPLICATION NO. PATENT NO. --------------______ 20031204 WO 2003-EP5529 20030527 WO 2003099268 A1 PΙ W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG 20020529 20031211 DE 2002-10224107 DE 10224107 A1 . AU 2003-240717 EP 2003-730120 20031212 20030527 A1 AU 2003240717 20050223 ` 20030527 A1 EP 1507520 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK US 2004-998164 20041129 A1 20050623 US 2005137194 US 2005-545901 20050817 Al 20060803 US 2006168942 20020529 Α PRAI DE 2002-10224107 W 20030527 WO 2003-EP5529 OS MARPAT 140:13084 The invention discloses the use of a combination of opioids (e.g. AB tramadol) with other active substances for producing a drug for the treatment of urinary urgency or urinary incontinence. The invention also relates to corresponding medicaments and to a method for treating urinary urgency or urinary incontinence. THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

---Logging off of STN---

Executing the logoff script...

=> LOG Y

SINCE FILE TOTAL COST IN U.S. DOLLARS ENTRY SESSION 27.99 29.56 FULL ESTIMATED COST DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE TOTAL ENTRY SESSION -6.75 -6.75 CA SUBSCRIBER PRICE

STN INTERNATIONAL LOGOFF AT 12:25:28 ON 02 NOV 2006

Connection closed by remote host

Connecting via Winsock to STN

Welcome to STN International! Enter x:x

LOGINID: ssptalxn1621

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

NEWS 1 Web Page URLs for STN Seminar Schedule - N. America
NEWS 2 "Ask CAS" for self-help around the clock
NEWS 3 AUG 09 INSPEC enhanced with 1898-1968 archive
NEWS 4 AUG 28 ADISCTI Reloaded and Enhanced
NEWS 5 AUG 30 CA(SM)/CAplus(SM) Austrian patent law changes
NEWS 6 SEP 11 CA/CAplus enhanced with more pre-1907 records
NEWS 7 SEP 21 CA/CAplus fields enhanced with simultaneous left and right truncation

NEWS 8 SEP 25 CA(SM)/CAplus(SM) display of CA Lexicon enhanced

NEWS 9 SEP 25 CAS REGISTRY(SM) no longer includes Concord 3D coordinates NEWS 10 SEP 25 CAS REGISTRY(SM) updated with amino acid codes for pyrrolysine

NEWS 11 SEP 28 CEABA-VTB classification code fields reloaded with new classification scheme

NEWS 12 OCT 19 LOGOFF HOLD duration extended to 120 minutes

NEWS 13 OCT 19 E-mail format enhanced

NEWS 14 OCT 23 Option to turn off MARPAT highlighting enhancements available

NEWS 15 OCT 23 CAS Registry Number crossover limit increased to 300,000 in multiple databases

NEWS 16 OCT 23 The Derwent World Patents Index suite of databases on STN has been enhanced and reloaded

NEWS 17 OCT 30 CHEMLIST enhanced with new search and display field

NEWS EXPRESS JUNE 30 CURRENT WINDOWS VERSION IS V8.01b, CURRENT MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP), AND CURRENT DISCOVER FILE IS DATED 26 JUNE 2006.

NEWS HOURS STN Operating Hours Plus Help Desk Availability
NEWS LOGIN Welcome Banner and News Items

NEWS IPC8 For general information regarding STN implementation of IPC 8 NEWS X25 X.25 communication option no longer available

Enter NEWS followed by the item number or name to see news on that specific topic.

All use of STN is subject to the provisions of the STN Customer agreement. Please note that this agreement limits use to scientific research. Use for software development or design or implementation of commercial gateways or other similar uses is prohibited and may result in loss of user privileges and other penalties.

FILE 'HOME' ENTERED AT 12:32:57 ON 02 NOV 2006

=> file reg COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.21 0.21

FULL ESTIMATED COST

FILE 'REGISTRY' ENTERED AT 12:33:15 ON 02 NOV 2006 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2006 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 1 NOV 2006 HIGHEST RN 912260-33-4 DICTIONARY FILE UPDATES: 1 NOV 2006 HIGHEST RN 912260-33-4

New CAS Information Use Policies, enter HELP USAGETERMS for details.

TSCA INFORMATION NOW CURRENT THROUGH June 30, 2006

Please note that search-term pricing does apply when conducting SmartSELECT searches.

REGISTRY includes numerically searchable data for experimental and predicted properties as well as tags indicating availability of experimental property data in the original document. For information on property searching in REGISTRY, refer to:

http://www.cas.org/ONLINE/UG/regprops.html

=>

Uploading C:\Program Files\Stnexp\Queries\fesoterodine generic.str

chain nodes :

7 8 9 10 11 12 13 14 15 16 23 24 25 26 27 28 29 30 31 ring nodes:

1 2 3 4 5 6 17 18 19 20 21 22

chain bonds :

5-7 7-8 7-17 8-9 9-10 10-11 10-12 11-15 11-16 12-13 12-14 19-26 22-23 23-24 24-25 24-29 26-27 26-30 26-31 27-28

ring bonds :

1-2 1-6 2-3 3-4 4-5 5-6 17-18 17-22 18-19 19-20 20-21 21-22

exact/norm bonds :

9-10 10-11 10-12 22-23 23-24 24-25 26-27

exact bonds :

5-7 7-8 7-17 8-9 11-15 11-16 12-13 12-14 19-26 24-29 26-30 26-31 27-28 normalized bonds:

1-2 1-6 2-3 3-4 4-5 5-6 17-18 17-22 18-19 19-20 20-21 21-22

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:CLASS 8:CLASS 9:CLASS 10:CLASS 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:CLASS 17:Atom 18:Atom 19:Atom 20:Atom 21:Atom 22:Atom 23:CLASS 24:CLASS 25:CLASS 26:CLASS

27:CLASS 28:CLASS 29:CLASS 30:CLASS 31:CLASS

L1

=> d L1

L1 HAS NO ANSWERS

L1

STR

Structure attributes must be viewed using STN Express guery preparation.

=> s L1

SAMPLE SEARCH INITIATED 12:33:38 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 5 TO ITERATE

100.0% PROCESSED

5 ITERATIONS

0 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS:

ONLINE **COMPLETE**

BATCH

COMPLETE

PROJECTED ITERATIONS: PROJECTED ANSWERS:

5 TO

0 TO 0

234

L2

0 SEA SSS SAM L1

=> file caplus

COST IN U.S. DOLLARS

SINCE FILE TOTAL

ENTRY SESSION

FULL ESTIMATED COST

0.44

0.65

FILE 'CAPLUS' ENTERED AT 12:33:44 ON 02 NOV 2006
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
COPYRIGHT (C) 2006 AMERICAN CHEMICAL SOCIETY (ACS)

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the American Chemical Society and is provided to assist you in searching

databases on STN. Any dissemination, distribution, copying, or storing of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 2 Nov 2006 VOL 145 ISS 19 FILE LAST UPDATED: 1 Nov 2006 (20061101/ED)

Effective October 17, 2005, revised CAS Information Use Policies apply. They are available for your review at:

http://www.cas.org/infopolicy.html

=> s L2

L3 0 L2

=>

---Logging off of STN---

Executing the logoff script...

=> LOG Y

COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.46 1.11

FULL ESTIMATED COST

STN INTERNATIONAL LOGOFF AT 12:34:30 ON 02 NOV 2006

Connecting via Winsock to STN

Welcome to STN International! Enter x:x

LOGINID:ssptalxn1621

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

Welcome to STN International

Web Page URLs for STN Seminar Schedule - N. America NEWS "Ask CAS" for self-help around the clock NEWS

INSPEC enhanced with 1898-1968 archive AUG 09 NEWS ADISCTI Reloaded and Enhanced AUG 28 NEWS

CA(SM)/CAplus(SM) Austrian patent law changes NEWS AUG 30

CA/CAplus enhanced with more pre-1907 records NEWS 6 SEP 11

CA/CAplus fields enhanced with simultaneous left and right SEP 21 NEWS truncation

CA(SM)/CAplus(SM) display of CA Lexicon enhanced SEP 25 NEWS 8

CAS REGISTRY(SM) no longer includes Concord 3D coordinates SEP 25 NEWS 9 .

CAS REGISTRY(SM) updated with amino acid codes for pyrrolysine SEP 25 NEWS 10

CEABA-VTB classification code fields reloaded with new SEP 28 NEWS 11 classification scheme

LOGOFF HOLD duration extended to 120 minutes OCT 19 NEWS 12

E-mail format enhanced OCT 19

NEWS 14 OCT 23 Option to turn off MARPAT highlighting enhancements available

NEWS 15 OCT 23 CAS Registry Number crossover limit increased to 300,000 in multiple databases

NEWS 16 OCT 23 The Derwent World Patents Index suite of databases on STN has been enhanced and reloaded

NEWS 17 OCT 30 CHEMLIST enhanced with new search and display field

NEWS EXPRESS JUNE 30 CURRENT WINDOWS VERSION IS V8.01b, CURRENT MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP), AND CURRENT DISCOVER FILE IS DATED 26 JUNE 2006.

NEWS HOURS STN Operating Hours Plus Help Desk Availability

NEWS LOGIN Welcome Banner and News Items

NEWS IPC8 For general information regarding STN implementation of IPC 8

NEWS X25 X.25 communication option no longer available

Enter NEWS followed by the item number or name to see news on that specific topic.

All use of STN is subject to the provisions of the STN Customer agreement. Please note that this agreement limits use to scientific research. Use for software development or design or implementation of commercial gateways or other similar uses is prohibited and may result in loss of user privileges and other penalties.

FILE 'HOME' ENTERED AT 14:38:17 ON 02 NOV 2006

=> file reg
COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.21 0.21

FULL ESTIMATED COST

FILE 'REGISTRY' ENTERED AT 14:38:27 ON 02 NOV 2006 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2006 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 1 NOV 2006 HIGHEST RN 912260-33-4 DICTIONARY FILE UPDATES: 1 NOV 2006 HIGHEST RN 912260-33-4

New CAS Information Use Policies, enter HELP USAGETERMS for details.

TSCA INFORMATION NOW CURRENT THROUGH June 30, 2006

Please note that search-term pricing does apply when conducting SmartSELECT searches.

REGISTRY includes numerically searchable data for experimental and predicted properties as well as tags indicating availability of experimental property data in the original document. For information on property searching in REGISTRY, refer to:

http://www.cas.org/ONLINE/UG/regprops.html

Uploading C:\Program Files\Stnexp\Queries\10532836.str

chain nodes :

7 8 9 10 11 12 13 14 15 16 17 18 19 20 21 22 23 24 25 26

ring nodes :

1 2 3 4 5 6

chain bonds :

5-7 7-8 7-17 8-9 9-10 10-11 10-14 11-12 11-13 14-15 14-16 17-18 17-22

18-19 19-20 19-21 22-23 22-25 22-26 23-24

ring bonds :

1-2 1-6 2-3 3-4 4-5 5-6

exact/norm bonds :

10-11 10-14 18-19 19-21 22-23

exact bonds :

5-7 7-8 7-17 8-9 9-10 11-12 11-13 14-15 14-16 17-18 17-22 19-20 22-25

22-26 23-24

normalized bonds :

1-2 1-6 2-3 3-4 4-5 5-6

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:CLASS 8:CLASS 9:CLASS 10:CLASS 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:CLASS 17:CLASS 18:CLASS 19:CLASS 20:CLASS 21:CLASS 22:CLASS 23:CLASS 24:CLASS 25:CLASS 26:CLASS

L1 STRUCTURE UPLOADED

=> d L1

L1 HAS NO ANSWERS

L1 STR

Structure attributes must be viewed using STN Express query preparation.

=> s L1 full

FULL SEARCH INITIATED 14:38:53 FILE 'REGISTRY'

FULL SCREEN SEARCH COMPLETED - 0 TO ITERATE

100.0% PROCESSED 0 ITERATIONS 0 ANSWERS

SEARCH TIME: 00.00.01

L2 0 SEA SSS FUL L1

=> file caplus

COST IN U.S. DOLLARS SINCE FILE TOTAL
ENTRY SESSION
FULL ESTIMATED COST 166.94 167.15

FILE 'CAPLUS' ENTERED AT 14:39:05 ON 02 NOV 2006 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
COPYRIGHT (C) 2006 AMERICAN CHEMICAL SOCIETY (ACS)

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the American Chemical Society and is provided to assist you in searching databases on STN. Any dissemination, distribution, copying, or storing of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 2 Nov 2006 VOL 145 ISS 19 FILE LAST UPDATED: 1 Nov 2006 (20061101/ED)

Effective October 17, 2005, revised CAS Information Use Policies apply. They are available for your review at:

http://www.cas.org/infopolicy.html

=> s L2

L3

0 L2

---Logging off of STN---

=>

Executing the logoff script...

=> LOG Y

COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.46 167.61

FULL ESTIMATED COST

STN INTERNATIONAL LOGOFF AT 14:39:22 ON 02 NOV 2006

Connecting via Winsock to STN

Welcome to STN International! Enter x:x

LOGINID: ssptalxn1621

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

NEWS 1 Web Page URLs for STN Seminar Schedule - N. America

NEWS 2 "Ask CAS" for self-help around the clock

NEWS 3 AUG 09 INSPEC enhanced with 1898-1968 archive

NEWS 4 AUG 28 ADISCTI Reloaded and Enhanced

NEWS 5 AUG 30 CA(SM)/CAplus(SM) Austrian patent law changes

NEWS 6 SEP 11 CA/CAplus enhanced with more pre-1907 records

NEWS 7 SEP 21 CA/Caplus fields enhanced with simultaneous left and right truncation

NEWS 8 SEP 25 CA(SM)/CAplus(SM) display of CA Lexicon enhanced

NEWS 9 SEP 25 CAS REGISTRY(SM) no longer includes Concord 3D coordinates

NEWS 10 SEP 25 CAS REGISTRY(SM) updated with amino acid codes for pyrrolysine

NEWS 11 SEP 28 CEABA-VTB classification code fields reloaded with new classification scheme

NEWS 12 OCT 19 LOGOFF HOLD duration extended to 120 minutes

NEWS 13 OCT 19 E-mail format enhanced

NEWS 14 OCT 23 Option to turn off MARPAT highlighting enhancements available

NEWS 15 OCT 23 CAS Registry Number crossover limit increased to 300,000 in multiple databases

NEWS 16 OCT 23 The Derwent World Patents Index suite of databases on STN has been enhanced and reloaded

NEWS 17 OCT 30 CHEMLIST enhanced with new search and display field

NEWS EXPRESS JUNE 30 CURRENT WINDOWS VERSION IS V8.01b, CURRENT MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP), AND CURRENT DISCOVER FILE IS DATED 26 JUNE 2006.

NEWS HOURS STN Operating Hours Plus Help Desk Availability

NEWS LOGIN Welcome Banner and News Items

NEWS IPC8 For general information regarding STN implementation of IPC 8

NEWS X25 X.25 communication option no longer available

Enter NEWS followed by the item number or name to see news on that specific topic.

All use of STN is subject to the provisions of the STN Customer agreement. Please note that this agreement limits use to scientific research. Use for software development or design or implementation of commercial gateways or other similar uses is prohibited and may result in loss of user privileges and other penalties.

FILE 'HOME' ENTERED AT 14:39:50 ON 02 NOV 2006

=> file reg
COST IN U.S. DOLLARS

FULL ESTIMATED COST

SINCE FILE TOTAL ENTRY SESSION 0.21 0.21

FILE 'REGISTRY' ENTERED AT 14:40:01 ON 02 NOV 2006
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
COPYRIGHT (C) 2006 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 1 NOV 2006 HIGHEST RN 912260-33-4 DICTIONARY FILE UPDATES: 1 NOV 2006 HIGHEST RN 912260-33-4

New CAS Information Use Policies, enter HELP USAGETERMS for details.

TSCA INFORMATION NOW CURRENT THROUGH June 30, 2006

Please note that search-term pricing does apply when conducting SmartSELECT searches.

REGISTRY includes numerically searchable data for experimental and predicted properties as well as tags indicating availability of experimental property data in the original document. For information on property searching in REGISTRY, refer to:

http://www.cas.org/ONLINE/UG/regprops.html

=> s fesoterodine

L1 2 FESOTERODINE

=> d fcn

L1 ANSWER 1 OF 2 REGISTRY COPYRIGHT 2006 ACS on STN

Propanoic acid, 2-methyl-, 2-[(1R)-3-[bis(1-methylethyl)amino]-1-phenylpropyl]-4-(hydroxymethyl)phenyl ester, (2E)-2-butenedioate (1:1) (salt) (9CI) (CA INDEX NAME)

OTHER NAMES:

CN Fesoterodine fumarate

CN SMP 8272

CN SPM 907

=> d L1

L1 ANSWER 1 OF 2 REGISTRY COPYRIGHT 2006 ACS on STN

RN 286930-03-8 REGISTRY

ED Entered STN: 21 Aug 2000

CN Propanoic acid, 2-methyl-, 2-[(1R)-3-[bis(1-methylethyl)amino]-1-

phenylpropyl]-4-(hydroxymethyl)phenyl ester, (2E)-2-butenedioate (1:1) (salt) (9CI) (CA INDEX NAME) OTHER NAMES: Fesoterodine fumarate CN SMP 8272 CN SPM 907 CN STEREOSEARCH FS C26 H37 N O3 . C4 H4 O4 MF SR ADISINSIGHT, BIOSIS, CA, CAPLUS, CBNB, IMSDRUGNEWS, LC STN Files: IMSPATENTS, IMSRESEARCH, PHAR, PROUSDDR, RTECS*, SYNTHLINE, TOXCENTER, USAN, USPATFULL (*File contains numerically searchable property data) CM

CRN 286930-02-7 CMF C26 H37 N O3

Absolute stereochemistry. Rotation (+).

CM 2

CRN 110-17-8 CMF C4 H4 O4

Double bond geometry as shown.

2 REFERENCES IN FILE CA (1907 TO DATE)

2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

=> s fesoterodine isobutyrate

2 FESOTERODINE

1530 ISOBUTYRATE

O FESOTERODINE ISOBUTYRATE

(FESOTERODINE (W) ISOBUTYRATE)

=> s fesoteridine derivatives

O FESOTERIDINE

165 DERIVATIVES

L3 0 FESOTERIDINE DERIVATIVES

(FESOTERIDINE (W) DERIVATIVES)

=> file caplus

L2

COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 28.92 29.13

FULL ESTIMATED COST

FILE 'CAPLUS' ENTERED AT 14:41:33 ON 02 NOV 2006
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
COPYRIGHT (C) 2006 AMERICAN CHEMICAL SOCIETY (ACS)

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the American Chemical Society and is provided to assist you in searching databases on STN. Any dissemination, distribution, copying, or storing of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 2 Nov 2006 VOL 145 ISS 19 FILE LAST UPDATED: 1 Nov 2006 (20061101/ED)

Effective October 17, 2005, revised CAS Information Use Policies apply. They are available for your review at:

http://www.cas.org/infopolicy.html

=> s fesoteridine

0 FESOTERIDINE

L4 0 FESOTERIDINE

=> s fesoterodine

L5 9 FESOTERODINE

=> d L5 1-9 all

- L5 ANSWER 1 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
- AN 2006:1133705 CAPLUS
- ED Entered STN: 30 Oct 2006
- TI Treatment of the overactive bladder syndrome with muscarinic receptor antagonists a matter of metabolites?
- AU Michel, Martin C.; Hegde, Sharath S.
- CS Department of Pharmacology & Pharmacotherapy, Academic Medical Center, University of Amsterdam, Meibergdreef 15, Amsterdam, 1105 AZ, Neth.
- SO Naunyn-Schmiedeberg's Archives of Pharmacology (2006), 374(2), 79-85 CODEN: NSAPCC; ISSN: 0028-1298
- PB Springer
- DT Journal
- LA English
- CC 1 (Pharmacology)
- Antagonists of muscarinic acetylcholine receptors, such as darifenacin, oxybutynin, propiverine, solifenacin, tolterodine, and trospium, are the mainstay of the treatment of the overactive bladder syndrome. Fesoterodine is a newer drug awaiting regulatory approval. We briefly review the pharmacol. activity of their metabolites and discuss how active metabolites may contribute to their efficacy and tolerability in vivo. Except for trospium, and perhaps solifenacin, all of the above drugs form active metabolites, and their presence and activity need to be taken into consideration when elucidating relationships between pharmacokinetics and pharmacodynamics of these drugs. Moreover, the ratios between parent compds. and metabolites may differ depending on genotype of the metabolizing enzymes, concomitant medication, and/or drug formulation. Differential generation of active metabolites of darifenacin or tolterodine are unlikely to influence the overall clin. profile of

these drugs in a major way because the active metabolites exhibit a similar pharmacol. profile as the parent compound In contrast, metabolites of oxybutynin and propiverine may behave quant. or even qual. differently from their parent compds. and this may have an impact on the overall clin. profile of these drugs. We conclude that more comprehensive studies of drug metabolites are required for an improved understanding of their clin. effects.

```
ANSWER 2 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
L5
    2006:76147 CAPLUS
AN
    144:156740
DN
    Entered STN: 27 Jan 2006
ED
    Combinations of statins with bronchodilators for treatment of respiratory
TI
    disorders
    Lindmark, Bertil; Thoren, Anders Ingemar
IN
    AstraZeneca AB, Swed.; AstraZeneca UK Limited
PΑ
SO
    PCT Int. Appl., 18 pp.
    CODEN: PIXXD2
DT
    Patent
    English
LΑ
IC
    ICM A61K031-40
        A61K031-505; A61K031-58; A61K031-165; A61P011-00; A61P011-06;
         A61P011-08
CC
    63-6 (Pharmaceuticals)
FAN.CNT 1
                              DATE
                                          APPLICATION NO.
                                                                DATE
    PATENT NO.
                       KIND
                                          ______
                              _____
     _____
                       ____
                              20060126 WO 2005-GB2413
    WO 2006008437
                                                                20050620
                        A1
PI
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
            CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
            GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ,
            LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA,
            NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK,
            SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU,
            ZA, ZM, ZW
        RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE,
            IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF,
            CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM,
            KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG,
            KZ, MD, RU, TJ, TM
                               20040715
PRAI GB 2004-15789
                         Α
CLASS
                CLASS PATENT FAMILY CLASSIFICATION CODES
 PATENT NO.
                      ______
 -----
                _ _ _ _
WO 2006008437
                ICM
                       A61K031-40
                       A61K031-505; A61K031-58; A61K031-165; A61P011-00;
                ICS
                       A61P011-06; A61P011-08
                       A61K0031-40 [ICM,7]; A61K0031-505 [ICS,7]; A61K0031-58
                IPCI
                       [ICS,7]; A61K0031-165 [ICS,7]; A61P0011-00 [ICS,7];
                       A61P0011-06 [ICS,7]; A61P0011-08 [ICS,7]
    The invention provides medicaments comprising combinations of
AΒ
    bronchodilators, glucocorticosteroids and HMG-CoA reductase inhibitors in
    the treatment of respiratory disorders such as chronic obstructive
    pulmonary disease (COPD). For example, a metered dose inhaler contained
    per dose formoterol fumarate dihydrate 4.5 μg, budesonide 160 μg,
    rosuvastatin 1 mg, and HFA 227 50 µL. Also, an inhalation/oral
    combination comprised an aerosol formulation containing per dose formoterol
    fumarate dihydrate 4.5 \mu g and budesonide 160 \mu g, and a tablet
    formulation containing rosuvastatin 10 mg.
    bronchodilator glucocorticosteroid statin respiratory disease; HMG CoA
ST
    reductase inhibitor bronchodilator respiratory disease
IT
    Drug delivery systems
        (aerosols, inhalants; combinations of statins with bronchodilators for
       treatment of respiratory disorders)
IT
    Lung, disease
```

```
(chronic obstructive pulmonary disease; combinations of statins with
        bronchodilators for treatment of respiratory disorders)
IT
     Bronchodilators
     Cholinergic antagonists
     Combination chemotherapy
     Respiratory system, disease
     β2-Adrenoceptor agonists
        (combinations of statins with bronchodilators for treatment of
        respiratory disorders)
     Glucocorticoids
IT
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (combinations of statins with bronchodilators for treatment of
       respiratory disorders)
    Drug delivery systems
IT
        (inhalants; combinations of statins with bronchodilators for treatment
       of respiratory disorders)
    Drug delivery systems
IT
        (powders, inhalants; combinations of statins with bronchodilators for
        treatment of respiratory disorders)
TT
    Drug delivery systems
        (tablets; combinations of statins with bronchodilators for treatment of
       respiratory disorders)
                            53-03-2, Prednisone
                                                   100-76-5D, Quinuclidine,
TΤ
     50-24-8, Prednisolone
                                                   3385-03-3, Flunisolide
     derivs. 124-94-7, Triamcinolone 596-51-0
                                25990-43-6, Mepenzolate
     4419-39-0, Beclomethasone
                                                           51333-22-3,
                  60135-22-0, Flumoxonide
                                            60205-81-4, Ipratropium
     Budesonide
                                                       75330-75-5, Lovastatin
     73573-87-2, Formoterol
                             73573-88-3, Mevastatin
     79902-63-9, Simvastatin 81093-37-0, Pravastatin 81732-65-2, Bambuterol
     85197-77-9, Tipredane 89365-50-4, Salmeterol 90566-53-3, Fluticasone
     93957-54-1, Fluvastatin 99571-64-9, Oxitropium 105102-22-5, Mometasone
     120815-74-9, Butixocort 124937-51-5, Tolterodine
                                                          126544-47-6,
                                              133099-04-4, Darifenacin
                 129260-79-3, Loteprednol
     Ciclesonide
                                136310-93-5, Tiotropium bromide
                                                                   137888-11-0,
     134523-00-5, Atorvastatin
              144459-70-1, Rofleponide
                                         145599-86-6, Cerivastatin
    TA 2005
                                                           183814-30-4,
                                182069-13-2, ETIPREDNOL
     170105-16-5, Imidafenacin
                                     186691-13-4, Tiotropium
                                                               192056-79-4
     Formoterol fumarate dihydrate
                               286930-02-7, Fesoterodine
     242478-37-1, Solifenacin
                                397864-44-7, 6α,9α-Difluoro-
     287714-41-4, Rosuvastatin
     17\alpha-[(2-furanylcarbonyl)oxy]-11\beta-hydroxy-16\alpha-methyl-3-oxo-
     androsta-1,4-diene-17β-carbothioic acid S-fluoromethyl ester
                  452339-68-3, 3-[4-[[6-[[(2R)-2-Hydroxy-2-[4-hydroxy-3-
     398455-25-9
     (hydroxymethyl)phenyl]ethyl]amino]hexyl]oxy]butyl]benzenesulfonamide
                                            867022-63-7
                 678160-57-1, Zoticasone
     463934-65-8
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (combinations of statins with bronchodilators for treatment of
        respiratory disorders)
     9028-35-7, HMG-CoA reductase
IT
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (inhibitors, statins; combinations of statins with bronchodilators for
        treatment of respiratory disorders)
TT
     147511-69-1
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (pitavastatin; combinations of statins with bronchodilators for
        treatment of respiratory disorders)
              THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
RE
(1) Harlan, J; WO 0048626 A 2000 CAPLUS
(2) Kao, P; US 2005119330 A1 2005
(3) Takeda Chemical Industries Ltd; EP 1275388 A 2003 CAPLUS
    ANSWER 3 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
L5
ΑN
    2004:878361 CAPLUS
DN
    141:370546
                  22 Oct 2004
ED
    Entered STN:
    Highly pure bases of 3,3-diphenyl propylamine monoesters for use in
TI
```

```
transdermal delivery systems
    Breitenbach, Armin; Meese, Claus; Wolff, Hans-Michael; Drews, Roland
IN
    Schwarz Pharma Ag, Germany
PA
    PCT Int. Appl., 72 pp.
SO
    CODEN: PIXXD2
DT
    Patent
    German
LΑ
    ICM C07C217-62
IC
    ICS A61K031-135; C07C213-10; A61P013-00
    63-6 (Pharmaceuticals)
CC
    Section cross-reference(s): 1
FAN.CNT 1
                                         APPLICATION NO.
    PATENT NO.
                              DATE
                                                                DATE
                        KIND
                             -----
    -----
                       ____
                                          ______
                             20041021 WO 2004-EP3567 20040403
    WO 2004089872
PΙ
                       A1
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
            CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
            GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
            NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
            TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
        RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
            BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,
            ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI,
            SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN,
            TD, TG
                                          DE 2003-10315917
                                                                20030408
                               20041118
    DE 10315917
                        A1
                                       AU 2004-228163 20040403
                       A1
                               20041021
    AU 2004228163
                                                               20040403
                                        CA 2004-2505848
    CA 2505848
                        AA
                               20041021
                                       BR 2004-6221
                                                                20040403
    BR 2004006221
                       Α
                               20050809
                                        EP 2004-725610
                                                                20040403
                              20060111
    EP 1613584
                        A1
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK,
                                                           20040403
                                        CN 2004-80009224
    CN 1802345
                              20060712
                       Α
                                                                20040403
                        T2
                                          JP 2006-504989
    JP 2006522758
                              20061005
                                                                20050426
                                          US 2005-532836
    NO 2005005078
                       A1 20060119
                                          NO 2005-5078
                                                                20051031
                       A 20051031
PRAI DE 2003-10315917
                             20030408
                       Α
                       W
                              20040403
    WO 2004-EP3567
CLASS
                CLASS PATENT FAMILY CLASSIFICATION CODES
PATENT NO.
------
               ----
                      ______
WO 2004089872
                ICM
                       C07C217-62
                       A61K031-135; C07C213-10; A61P013-00
                ICS
                       C07C0217-62 [ICM,7]; C07C0217-00 [ICM,7,C*];
                IPCI
                       A61K0031-135 [ICS,7]; C07C0213-10 [ICS,7]; C07C0213-00
                       [ICS,7,C*]; A61P0013-00 [ICS,7]
                       A61K0009-70 [I,C*]; A61K0009-70 [I,A]; A61K0031-135
                IPCR
                       [I,C*]; A61K0031-135 [I,A]; A61P0013-00 [I,C*];
                       A61P0013-00 [I,A]; C07C0213-00 [I,C*]; C07C0213-10
                       [I,A]; C07C0217-00 [I,C*]; C07C0217-62 [I,A]
                       A61K009/70E; A61K031/135; C07C213/10; C07C217/62
                ECLA
                       C07C0219-28 [ICM, 7]; C07C0219-00 [ICM, 7, C*]
DE 10315917
                IPCI
                IPCR
                       A61K0009-70 [I,C*]; A61K0009-70 [I,A]; A61K0031-135
                       [I,C*]; A61K0031-135 [I,A]; A61P0013-00 [I,C*];
                       A61P0013-00 [I,A]; C07C0213-00 [I,C*]; C07C0213-10
                       [I,A]; C07C0217-00 [I,C*]; C07C0217-62 [I,A]
                       A61K009/70E; A61K031/135; C07C213/10; C07C217/62
                ECLA
                       C07C0217-62 [ICM, 7]; C07C0217-00 [ICM, 7, C*];
AU 2004228163
                IPCI
                       A61K0031-135 [ICS,7]; C07C0213-10 [ICS,7]; C07C0213-00
                       [ICS,7,C*]; A61P0013-00 [ICS,7]
                       A61K0009-70 [I,C*]; A61K0009-70 [I,A]; A61K0031-135
                IPCR
                       [I,C*]; A61K0031-135 [I,A]; A61P0013-00 [I,C*];
                       A61P0013-00 [I,A]; C07C0213-00 [I,C*]; C07C0213-10
                       [I,A]; C07C0217-00 [I,C*]; C07C0217-62 [I,A]
```

```
CA 2505848
                 IPCI
                        C07C0217-62 [ICM,7]; C07C0217-00 [ICM,7,C*];
                        A61P0013-00 [ICS,7]; C07C0213-10 [ICS,7]; C07C0213-00
                        [ICS,7,C*]; A61K0031-135 [ICS,7]
                        A61K0009-70 [I,C*]; A61K0009-70 [I,A]; A61K0031-135
                 IPCR
                        [I,C*]; A61K0031-135 [I,A]; A61P0013-00 [I,C*];
                        A61P0013-00 [I,A]; C07C0213-00 [I,C*]; C07C0213-10
                        [I,A]; C07C0217-00 [I,C*]; C07C0217-62 [I,A]
                        A61K009/70E; A61K031/135; C07C213/10; C07C217/62
                 ECLA
                        C07C0217-62 [ICM,7]; C07C0217-00 [ICM,7,C*];
 BR 2004006221
                 IPCI
                        A61K0031-135 [ICS,7]; C07C0213-10 [ICS,7]; C07C0213-00
                        [ICS,7,C*]; A61P0013-00 [ICS,7]
                        A61K0009-70 [I,C*]; A61K0009-70 [I,A]; A61K0031-135
                 IPCR
                        [I,C*]; A61K0031-135 [I,A]; A61P0013-00 [I,C*];
                        A61P0013-00 [I,A]; C07C0213-00 [I,C*]; C07C0213-10
                        [I,A]; C07C0217-00 [I,C*]; C07C0217-62 [I,A]
                        A61K009/70E; A61K031/135; C07C213/10; C07C217/62
                 ECLA
                        C07C0217-62 [ICM,7]; C07C0217-00 [ICM,7,C*];
 EP 1613584
                 IPCI
                        A61K0031-135 [ICS,7]; C07C0213-10 [ICS,7]; C07C0213-00
                        [ICS,7,C*]; A61P0013-00 [ICS,7]
                        A61K0009-70 [I,C*]; A61K0009-70 [I,A]; A61K0031-135
                 IPCR
                        [I,C*]; A61K0031-135 [I,A]; A61P0013-00 [I,C*];
                        A61P0013-00 [I,A]; C07C0213-00 [I,C*]; C07C0213-10
                        [I,A]; C07C0217-00 [I,C*]; C07C0217-62 [I,A]
                        A61K009/70E; A61K031/135; C07C213/10; C07C217/62
                 ECLA
                        C07C0217-62 [I,A]; C07C0217-00 [I,C*]; A61K0031-135
 CN 1802345
                 IPCI
                        [I,A]; C07C0213-10 [I,A]; C07C0213-00 [I,C*];
                        A61P0013-00 [I,A]
                        A61K009/70E; A61K031/135; C07C213/10; C07C217/62
                 ECLA
 JP 2006522758
                 IPCI
                        C07C0219-28 [I,A]; C07C0219-00 [I,C*]; C07C0213-08
                        [I,A]; C07C0213-10 [I,A]; C07C0213-00 [I,C*];
                        A61K0031-222 [I,A]; A61K0031-21 [I,C*]; A61P0013-02
                        [I,A]; A61P0013-10 [I,A]; A61P0013-00 [I,C*];
                        A61K0009-70 [I,A]; A61K0047-32 [I,A]; C07B0053-00 [N,A]
                        4C076/AA74; 4C076/AA95; 4C076/CC17; 4C076/EE10M;
                 FTERM
                        4C076/EE10Q; 4C076/EE12M; 4C076/EE12Q; 4C076/EE13M;
                        4C076/EE13Q; 4C076/EE47M; 4C076/EE47Q; 4C076/EE48M;
                        4C076/EE48Q; 4C076/FF31; 4C076/FF63; 4C076/FF68;
                        4C206/AA01; 4C206/AA02; 4C206/DB02; 4C206/DB57;
                        4C206/KA13; 4C206/MA02; 4C206/MA05; 4C206/MA33;
                        4C206/MA36; 4C206/MA48; 4C206/MA52; 4C206/MA55;
                        4C206/MA56; 4C206/MA76; 4C206/MA83; 4C206/NA03;
                        4C206/NA12; 4C206/NA13; 4C206/ZA81; 4H006/AA01;
                        4H006/AA02; 4H006/AA03; 4H006/AB20; 4H006/AC52;
                        4H006/AC81; 4H006/AD16; 4H006/BB11; 4H006/BB12;
                        4H006/BB15; 4H006/BB16; 4H006/BB17; 4H006/BB31;
                        4H006/BC16; 4H006/BE12; 4H006/BE13; 4H006/BJ50;
                        4H006/BN10; 4H006/BT16; 4H006/BU36
                        C07C0229-52 [I,A]; C07C0229-00 [I,C*]; A61K0031-24
 US 2006014832
                 IPCI
                        [I,A]; A61K0031-21 [I,C*]
                        514/540.000; 560/136.000
                 NCL
                        A61K009/70E; A61K031/135; C07C213/10; C07C217/62
                 ECLA
                        C07C0217-62 [ICM,7]; C07C0217-00 [ICM,7,C*];
NO 2005005078
                 IPCI
                        A61K0031-135 [ICS,7]; A61K0009-70 [ICS,7]; A61P0013-00
                        [ICS, 7]
                        A61K0009-70 [I,C*]; A61K0009-70 [I,A]; A61K0031-135
                 IPCR
                        [I,C*]; A61K0031-135 [I,A]; A61P0013-00 [I,C*];
                        A61P0013-00 [I,A]; C07C0213-00 [I,C*]; C07C0213-10
                        [I,A]; C07C0217-00 [I,C*]; C07C0217-62 [I,A]
                        A61K009/70E; A61K031/135; C07C213/10; C07C217/62
                 ECLA
OS
    MARPAT 141:370546
```

GI

AB The invention relates to a compound of general formula (I) wherein A represents deuterium or hydrogen, R represents a group selected from C1-6 alkyl, C3-10 cycloalkyl or Ph, which can be substituted by C1-3 alkoxy, fluorine, chlorine, bromine, iodine, nitro, amino, hydroxyl, oxo, mercapto or deuterium. The C atom marked with a \star (star) can be present in an (R) configuration, in an (S)-configuration or a mixture thereof. The invention is characterized in that the above-mentioned compds. are free bases with a degree of purity of more than 97 wt %. The invention also relates to a method for the production of highly pure compds. of general formula (I) and to the use thereof in the production of medicaments. Thus (R)-2-[3-(Diisopropylamino) -1-phenylpropyl] -4-(hydroxymethyl)phenol was reacted with isobutyric acid chloride to form fesoterodine. Fesoterodine was purified via the formation of its fumaric acid salt. 1.5 G of the highly pure fesoterodine was mixed with 8.5 q silicone adhesive Bio-PSA 7-4300 and applied to a foil in order to prepare a transdermal delivery system.

ST fesoterodine purifn monoester transdermal delivery system

IT Ion exchangers

(basic; highly pure bases of 3,3-di-Ph propylamine monoesters for use in transdermal delivery systems)

IT Bladder

(detrusor muscle, hyperactivity of; highly pure bases of 3,3-di-Ph propylamine monoesters for use in transdermal delivery systems)

IT Adhesives

Chirality

Crystallization

Dissolution

(highly pure bases of 3,3-di-Ph propylamine monoesters for use in transdermal delivery systems)

IT Amines, reactions.

Bicarbonates

RL: RCT (Reactant); RACT (Reactant or reagent)

(highly pure bases of 3,3-di-Ph propylamine monoesters for use in transdermal delivery systems)

IT Bladder, disease

(incontinence; highly pure bases of 3,3-di-Ph propylamine monoesters for use in transdermal delivery systems)

IT Urinary system, disease

(nocturia; highly pure bases of 3,3-di-Ph propylamine monoesters for use in transdermal delivery systems)

IT Bladder, disease

(pollakisuria; highly pure bases of 3,3-di-Ph propylamine monoesters for use in transdermal delivery systems)

IT Amines, reactions

RL: RCT (Reactant); RACT (Reactant or reagent)

(polyamines, nonpolymeric; highly pure bases of 3,3-di-Ph propylamine monoesters for use in transdermal delivery systems)

```
Drug delivery systems
IT
        (transdermal; highly pure bases of 3,3-di-Ph propylamine monoesters for
       use in transdermal delivery systems)
    Drug delivery systems
IT
        (transmucosal; highly pure bases of 3,3-di-Ph propylamine monoesters
        for use in transdermal delivery systems)
     60-29-7, Diethyl ether, uses
                                   75-09-2, Dichloromethane, uses
IT
    Ethylmethylketone, uses 108-88-3, Toluene, uses
                                                        141-78-6,
                        1634-04-4, tert. Butylmethyl ether
    Ethylacetate, uses
    RL: NUU (Other use, unclassified); USES (Uses)
        (highly pure bases of 3,3-di-Ph propylamine monoesters for use in
        transdermal delivery systems)
  286930-02-7P, Fesoterodine
    RL: PRP (Properties); RCT (Reactant); SPN (Synthetic preparation); THU
     (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT
     (Reactant or reagent); USES (Uses)
        (highly pure bases of 3,3-di-Ph propylamine monoesters for use in
        transdermal delivery systems)
IT
     504415-91-2P, Bio-PSA 7-4300
    RL: PRP (Properties); SPN (Synthetic preparation); PREP (Preparation)
        (highly pure bases of 3,3-di-Ph propylamine monoesters for use in
        transdermal delivery systems)
IT
    777075-72-6P
    RL: PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use);
    BIOL (Biological study); PREP (Preparation); USES (Uses)
        (highly pure bases of 3,3-di-Ph propylamine monoesters for use in
        transdermal delivery systems)
    79-30-1, Isobutyric acid chloride
                                         110-17-8, Fumaric acid, reactions
IT
     207679-81-0
    RL: RCT (Reactant); RACT (Reactant or reagent)
        (highly pure bases of 3,3-di-Ph propylamine monoesters for use in
        transdermal delivery systems)
    5586-73-2D, 3,3-Diphenyl propylamine, monoesters
IT
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (highly pure bases of 3,3-di-Ph propylamine monoesters for use in
        transdermal delivery systems)
             THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
RE
(1) Kreilgaard, B; WO 0012070 A 2000 CAPLUS
(2) Nilv; WO 9411337 A 1994 CAPLUS
(3) Sanol Arznei Schwarz Gmbh; WO 9958478 A 1999 CAPLUS
(4) Sanol Arznei Schwarz Gmbh; WO 0135957 A 2001
    ANSWER 4 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
L5
    2004:875349 CAPLUS
ΑN
DN
    142:303234
    Entered STN: 22 Oct 2004
ED
    Mucosal adjuvants and delivery systems for oral and nasal vaccination
ΤI
    Baudner, Barbara C.; Verhoel, J. Coos; Junginger, Hans E.; del Giudice,
ΑU
    Giuseppe
CS
     IRIS Research Center, Siena, 53100, Italy
SO
    Drugs of the Future (2004), 29(7), 721-732
    CODEN: DRFUD4; ISSN: 0377-8282
PB
     Prous Science
     Journal; General Review
DT
LA
    English
CC
     63-0 (Pharmaceuticals)
     Section cross-reference(s): 15
    A review. The pillars of pharmacotherapy for overactive bladder (OAB) are
AΒ
     antimuscarinic agents which inhibit bladder smooth muscle contractions
     through interference with acetylcholine action on muscarinic receptors of
     the detrusor smooth muscle. Despite the availability of different
     antimuscarinic compds., physicians and patients remain dissatisfied with
     current treatments due to adverse events and/or insufficient efficacy.
```

Therefore, new agents with improved safety and efficacy profiles are

needed for a more effective treatment of overactive bladder. Fesoterodine is a novel bladder-selective muscarinic antagonist that has shown potent antimuscarinic activity in vitro and in vivo. In multiple investigations, the agent has been shown to be safe and well tolerated in subjects of different ethnic origin, age and gender; in poor and extensive CYP2D6 metabolizers; in subjects taking concomitant medication inhibiting CYP3A4; in fed or fasted states; and in those suffering from hepatic impairment. No clin. relevant changes in heart rate, blood pressure, ECG parameters or laboratory analyses have been seen with therapeutic doses of fesoterodine in these studies. Furthermore, in a phase II clin. trial in patients with OAB, fesoterodine demonstrated rapid and significant efficacy on a variety of endpoints. The results of this trial encouraged the manufacturer (SCHWARZ PHARMA) to initiate a phase III clin. trial program for fesoterodine.

- ST review mucosa adjuvant oral nasal vaccine
- IT Immunostimulants

(adjuvants; mucosal adjuvants and delivery systems for oral and nasal vaccination)

IT Muscarinic antagonists

Vaccines

(mucosal adjuvants and delivery systems for oral and nasal vaccination)

IT Drug delivery systems

(nasal; mucosal adjuvants and delivery systems for oral and nasal vaccination)

IT Drug delivery systems

(oral; mucosal adjuvants and delivery systems for oral and nasal vaccination)

- RE.CNT 169 THERE ARE 169 CITED REFERENCES AVAILABLE FOR THIS RECORD RE
- (1) Ackermann, L; Immunology 2004, V111, P75 CAPLUS
- (2) Agnello, D; J Clin Immunol 2003, V23, P147 CAPLUS
- (3) Allaoui-Attarki, K; Infect Immun 1997, V65, P853 CAPLUS
- (4) Andre, F; Vaccine 2003, V21, P593
- (5) Babiuk, L; Vaccine 2003, V21, P649 CAPLUS
- (6) Baca-Estrada, M; Vaccine 2000, V18, P2203 CAPLUS
- (7) Baldridge, J; Modern Vaccines Adjuvants Deliv Syst 2003
- (8) Baldridge, J; Vaccine 2000, V18, P2416 CAPLUS
- (9) Baudner, B; Infect Immun 2002, V70, P4785 CAPLUS
- (10) Baudner, B; J Infect Dis 2004, V189, P828 CAPLUS
- (11) Baudner, B; Vaccine 2003, V21, P3837 CAPLUS
- (12) Bienenstock, J; Bacterial Infections of Respiratory and Gastrointestinal Mucosae 1998, P9
- (13) Bienenstock, J; Lab Invest 1976, V35, P343 MEDLINE
- (14) Borchard, G; J Control Rel 1996, V39, P131 CAPLUS
- (15) Bouche, F; Vaccine 2003, V21, P2065 CAPLUS
- (16) Bowman, C; Infect Immun 2001, V69, P1528 CAPLUS
- (17) Boyaka, P; Adv Drug Deliv Rev 2001, V51, P71 CAPLUS
- (18) Boyaka, P; Immunol Res 1999, V20, P207 CAPLUS
- (19) Bradney, C; J Virol 2002, V76, P517 CAPLUS
- (20) Brandtzaeg, P; Nat Immunol 2001, V2, P1093 CAPLUS
- (21) Braun, M; J Exp Med 1999, V189, P541 CAPLUS
- (22) Buchan, G; Mol Immunol 2000, V379, P545
- (23) Cahill, E; Vaccine 1995, V13, P455 CAPLUS
- (24) Carcaboso, A; Intl J Pharm 2003, V260, P273 CAPLUS
- (25) Challacombe, S; Vaccine 1997, V15, P169 MEDLINE
- (26) Chikwamba, R; Transgenic Res 2002, V11, P479 CAPLUS
- (27) Childers, N; Infect Immun 1999, V67, P618 CAPLUS
- (28) Childers, N; Infect Immun 2000, V68, P5509 CAPLUS
- (29) Childers, N; J Dent Res 2002, V81, P48 MEDLINE
- (30) Choi, A; Vaccine 2002, V20, P1733 CAPLUS
- (31) Chun, S; J Immunol 1999, V163, P2393 CAPLUS
- (32) Cleland, J; Trends Biotechnol 1999, V17, P25 CAPLUS
- (33) Conway, M; Vaccine 2001, V19, P1940 CAPLUS
- (34) Coulter, A; Vaccine 2003, V21, P946 CAPLUS

```
(35) Czerkinsky, C; Immunol Rev 1999, V170, P197 CAPLUS
```

- (36) Davis, H; Vaccine 2000, V18, P1920 CAPLUS
- (37) Dehaan, L; Eur J Immunol 1998, V28, P1243 CAPLUS
- (38) Desai, M; Pharm Res 1997, V14, P1568 CAPLUS
- (39) Dickinson, B; Infect Immun 1995, V63, P1617 CAPLUS
- (40) Dodane, V; Intl J Pharm 1999, V182, P21 CAPLUS
- (41) Doherty, T; Infect Immun 2002, V70, P3111 CAPLUS
- (42) Duc Le, H; Infect Immun 2003, V71, P2810 MEDLINE
- (43) Dumais, N; J Infect Dis 2002, V186, P1098 CAPLUS
- (44) Dziekan, G; Bull World Health Org 2003, V81, P277
- (45) Ehreth, J; Vaccine 2003, V21, P596
- (46) Eldridge, J; J Control Rel. 1990, V11, P205 CAPLUS
- (47) Fagarasan, S; Nat Rev Immunol 2003, V3, P63 CAPLUS
- (48) Fagarasan, S; Nature 2001, V413, P639 CAPLUS
- (49) Finke, D; Curr Opin Genet Dev 2001, V11, P561 CAPLUS
- (50) Freidag, B; Infect Immun 2000, V68, P2948 CAPLUS
- (51) Furrie, E; Vaccine 2002, V20, P2254 CAPLUS
- (52) Gagliardi, M; Eur J Immunol 2000, V30, P2394 CAPLUS
- (53) Gallichan, W; J Immunol 2001, V166, P3451 CAPLUS
- (54) Gao, Y; World J Gastroenterol 2003, V9, P996 CAPLUS
- (55) Giuliani, M; J Exp Med 1998, V187, P1123 CAPLUS
- (56) Gupta, R; Dev Biol Stand 1998, V92, P63 CAPLUS
- (57) Han, M; J Vet Med Sci 1997, V59, P1109 CAPLUS
- (58) Hantman, M; Mucosal Immunology, 2nd Ed 1999, P759
- (59) Harokopakis, E; Infect Immun 1998, V66, P4299 CAPLUS
- (60) Hathaway, L; Cell Mol Life Sci 2000, V57, P323 CAPLUS
- (61) Heritage, P; Immunology 1998, V93, P249 CAPLUS
- (62) Herrmann, J; Virology 1999, V259, P148 CAPLUS
- (63) Hu, K; Adv Drug Deliv Rev 2001, V51, P149 CAPLUS
- (64) Hu, K; Clin Exp Immunol 1998, V113, P235 CAPLUS
- (65) Illum, L; Adv Drug Deliv Rev 2001, V51, P81 CAPLUS
- (66) Jabbal-Gill, I; Vaccine 1998, V16, P2039 CAPLUS
- (67) Jahnsen, F; Am J Respir Cell Mol Biol 2004, V30, P31 CAPLUS
- (68) Jakobsen, H; Infect Immun 2002, V70, P1443 CAPLUS
- (69) Jalava, K; Expert Rev Vaccines 2003, V2, P45 CAPLUS
- (70) Janeway, C; Annu Rev Immunol 2002, V20, P197 CAPLUS
- (71) Jepson, M; Adv Drug Deliv Rev 2004, V56, P511 CAPLUS
- (72) Johnson, M; Modern Vaccines Adjuvants Deliv Syst 2003
- (73) Jones, D; Infect Immun 1996, V64, P489 CAPLUS
- (74) Kato, Y; Curr Pharm Biotechnol 2003, V4, P303 CAPLUS
- (75) Kawasaki, K; Jpn J Infect Dis 2001, V54, P220 CAPLUS
- (76) Kotze, A; Pharm Res 1997, V14, P1197 CAPLUS
- (77) Kraehenbuhl, J; Annu Rev Cell Dev Biol 2000, V16, P301 CAPLUS
- (78) Krieg, A; Ann Rev Immunol 2002, V29, P709
- (79) Krieg, A; Antisense Nucleic Acid Drug Dev 2001, V11, P181 CAPLUS
- (80) Krieg, A; Nature 1995, V374, P546 CAPLUS
- (81) Kuklin, N; J Virol 1997, V71, P3138 CAPLUS
- (82) Larsen, D; Am J Vet Res 2002, V63, P653 CAPLUS
- (83) Levine, M; Bull Inst Pasteur 1995, V93, P243
- (84) Levine, M; Microbiol Rev 1983, V47, P510 CAPLUS
- (85) Levine, M; Vaccine 1999, V17, P22
- (86) Lundholm, P; Virus Res 2002, V82, P141 CAPLUS
- (87) Lycke, N; Cell Microbiol 2004, V6, P23 CAPLUS
- (88) Mahon, B; J Exp Med 1995, V181, P1285 CAPLUS
- (89) Marchetti, M; Vaccine 1998, V16, P33 CAPLUS
- (90) Marinaro, M; J Immunol 1999, V162, P114 CAPLUS
- (91) Mariotti, S; Vaccine 2002, V20, P2229 CAPLUS
- (92) McCluskie, M; J Immunol 1998, V161, P4463 CAPLUS
- (93) McCluskie, M; Vaccine 2000, V19, P413 CAPLUS
- (94) McNeela, E; Vaccine 2000, V19, P1188 CAPLUS
- (95) McNeela, E; Vaccine 2004, V22, P909 CAPLUS
- (96) Medzhitov, R; Nature 1997, V388, P394 CAPLUS
- (97) Mestecky, J; Mucosal Immunology, 2nd Ed 1999, P133
- (98) Michalek, S; Mucosal Immunology, 2nd Ed 1999, P759
- (99) Michie, C; Trends Mol Med 2002, V8, P6

```
(100) Mielcarek, N; Adv Drug Deliv Rev 2001, V51, P55 CAPLUS
```

- (101) Mills, K; Infect Immun 2003, V71, P726 CAPLUS
- (102) Moore, A; Vaccine 1999, V17, P2517 CAPLUS
- (103) Mor, G; J Clin Invest 1996, V98, P2700 CAPLUS
- (104) Mosmann, T; Immunol Today 1996, V17, P138 CAPLUS
- (105) Mossad, S; Cleve Clin J Med 2003, V70, P801
- (106) Murphy, T; New Engl J Med 2001, V344, P564 MEDLINE
- (107) Mutsch, M; New Engl J Med 2004, V350, P896 CAPLUS
- (108) Neutra, M; Annu Rev Immunol 1996, V14, P275 CAPLUS
- (109) Neutra, M; Cell 1996, V86, P345 CAPLUS
- (110) Neutra, M; Nat Immunol 2001, V2, P1004 CAPLUS
- (111) Ninomiya, A; Vaccine 2002, V19, P3123
- (112) Nishimura, K; Vaccine 1984, V2, P93 CAPLUS
- (113) Oggioni, M; Vaccine 2003, V21(Suppl 2), PS96
- (114) Okuda, K; Vaccine 2001, V19, P3681 CAPLUS
- (115) Olszewska, W; Infect Immun 2000, V68, P4923 CAPLUS
- (116) O'Hagan, D; Expert Rev Vaccines 2003, V2, P269 CAPLUS
- (117) O'Hagan, D; J Anat 1996, V189, P477 CAPLUS
- (118) O'Hagan, D; Novel Delivery Systems for Oral Vaccines 1994, P175 CAPLUS
- (119) Pasetti, M; Vaccine 2003, V21, P401 CAPLUS
- (120) Paul, W; STP Pharma Sci 2000, V10, P5 CAPLUS
- (121) Prince, G; J Virol 2003, V77, P13156 CAPLUS
- (122) Rappuoli, R; Immunol Today 1999, V20, P493 CAPLUS
- (123) Richards, C; J Infect Dis 1998, V177, P1451 CAPLUS
- (124) Roy, K; Nat Med 1999, V5, P387 CAPLUS
- (125) Ryan, E; Infect Immun 1999, V67, P6270 CAPLUS
- (126) Ryan, E; J Immunol 2000, V165, P5750 CAPLUS
- (127) Sala, F; Vaccine 2003, V21, P803 CAPLUS
- (128) Salkowski, C; Infect Immun 1997, V65, P3239 CAPLUS
- (129) Sanchez, J; Lancet 1994, V344, P1273 MEDLINE
- (130) Sato, Y; Science 1996, V273, P352 CAPLUS
- (131) Seo, J; Infect Immun 2002, V70, P1143 CAPLUS
- (132) Seong, S; Infect Immun 1999, V67, P3587 CAPLUS
- (133) Shreedhar, V; Infect Immun 2003, V71, P504 CAPLUS (134) Sieval, A; Carbohydr Polym 1998, V36, P157 CAPLUS
- (135) Singh, M; Vaccine 2001, V20, P594 CAPLUS
- (136) Smith, M; Exp Phys 1995, V80, P735 CAPLUS
- (137) Smith, R; Immunol Cell Biol 1998, V76, P263 CAPLUS
- (138) Smith, R; J Immunol 1999, V162, P5536 CAPLUS
- (139) Sommer, F; Infect Immun 2004, V72, P1029 CAPLUS
- (140) Staats, H; J Immunol 1999, V162, P6141 CAPLUS
- (141) Staats, H; J Immunol 2001, V167, P5386 CAPLUS
- (142) Steinman, R; J Clin Invest 2002, V109, P1519 CAPLUS
- (143) Streatfield, S; Expert Rev Vaccines 2003, V2, P763
- (144) Sun, J; J Immunol 1999, V163, P1045 CAPLUS
- (145) Tacket, C; J Infect Dis 2000, V182, P302 MEDLINE
- (146) Takahashi, I; J Infect Dis 1996, V173, P627 CAPLUS
- (147) Thanou, M; J Control Rel 2000, V64, P15 CAPLUS
- (148) Toka, F; J Virol 2003, V77, P12742 CAPLUS
- (149) Trach, D; Lancet 1997, V349, P231 MEDLINE
- (150) Underhill, D; Curr Opin Immunol 2002, V14, P103 CAPLUS
- (151) Vajdy, M; Adv Drug Deliv Rev 2001, V51, P127 CAPLUS
- (152) van Ginkel, F; J Immunol 2000, V165, P4778 CAPLUS
- (153) van der Lubben, I; STP Pharma Sci 2002, V12, P235 CAPLUS
- (154) van der Lubben, I; Vaccine 2003, V21, P1400 CAPLUS
- (155) Vasselon, T; Infect Immun 2002, V70, P1033 CAPLUS
- (156) Vega-Lopez, M; Modern Vaccines Adjuvants Deliv Syst 2003
- (157) Venkataprasad, N; Vaccine 1999, V17, P1814 CAPLUS
- (158) Verthelyi, D; J Immunol 2001, V166, P2372 CAPLUS
- (159) Verthelyi, D; J Immunol 2002, V168, P1659 CAPLUS
- (160) Wagner, H; Curr Opin Immunol 2002, V5, P62 CAPLUS
- (161) Warzecha, H; J Virol 2003, V77, P8702 CAPLUS
- (162) Weiner, H; Microbes Infect 2001, V3, P947 CAPLUS
- (163) Whittum-Hudson, J; Nat Med 1996, V2, P1116 CAPLUS (164) Wu, Y; Virology 2003, V313, P337 CAPLUS

```
(165) Xin, K; Immunology 1998, V94, P438 CAPLUS
(166) Xu, L; Clin Immunol 2000, V96, P205 CAPLUS
(167) Xu-Amano, J; Vaccine 1994, V12, P903 CAPLUS
(168) Yamamoto, M; J Infect Dis 2000, V182, P180 CAPLUS
(169) Yamanaka, T; Eur J Immunol 2001, V31, P107 CAPLUS
    ANSWER 5 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
L5
    2004:875348 CAPLUS
AN
    142:147630
DN
    Entered STN: 22 Oct 2004
ED
    Fesoterodine, an advanced antimuscarinic for the treatment of
TI
    overactive bladder: a safety update
ΑU
    Cole, Patrick
    Medical Information Dept., Prous Science, Barcelona, 08080, Spain
CS
    Drugs of the Future (2004), 29(7), 715-720
SO
    CODEN: DRFUD4; ISSN: 0377-8282
PB
    Prous Science
    Journal; General Review
DT
LΑ
    English
CC
    1-0 (Pharmacology)
    A review. The pillars of pharmacotherapy for overactive bladder (OAB) are
AB
    antimuscarinic agents which inhibit bladder smooth muscle contractions
    through interference with acetylcholine action on muscarinic receptors of
    the detrusor smooth muscle. Despite the availability of different
    antimuscarinic compds., physicians and patients remain dissatisfied with
    current treatments due to adverse events and/or insufficient efficacy.
    Therefore, new agents with improved safety and efficacy profiles are
    needed for a more effective treatment of overactive bladder.
    Fesoterodine is a novel bladder-selective muscarinic antagonist
    that has shown potent antimuscarinic activity in vitro and in vivo.
    multiple investigations, the agent has been shown to be safe and well
    tolerated in subjects of different ethnic origin, age and gender; in poor
    and extensive CYP2D6 metabolizers; in subjects taking concomitant
    medication inhibiting CYP3A4; in fed or fasted states; and in those
    suffering from hepatic impairment. No clin. relevant changes in heart
    rate, blood pressure, ECG parameters or laboratory analyses have been seen with
    therapeutic doses of fesoterodine in these studies.
    Furthermore, in a phase II clin. trial in patients with OAB,
    fesoterodine demonstrated rapid and significant efficacy on a
    variety of endpoints. The results of this trial encouraged the
    manufacturer (SCHWARZ PHARMA) to initiate a phase III clin. trial program
    for fesoterodine.
    review fesoterodine antimuscarinic overactive bladder
ST
    Combination chemotherapy
    Drug interactions
    Human
    Muscarinic antagonists
        (advanced antimuscarinic fesoterodine for treatment of
       overactive bladder)
    Bladder, disease
IT
        (hyperreflexia; advanced antimuscarinic fesoterodine for
       treatment of overactive bladder)
    286930-02-7, Fesoterodine
TT
    RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological
    activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (advanced antimuscarinic fesoterodine for treatment of
       overactive bladder)
             THERE ARE 24 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
(1) Abrams, P; Urology 2003, V61, P37
(2) Andersson, K; BJU Int 1999, V84, P923 CAPLUS
(3) Andersson, K; Bailliere's Best Pract Res Clin Obstet Gynaecol 2000, V14,
```

P291 MEDLINE

(4) Anon; Data on file Schwarz BioSciences

(5) Anon; ICS/IUGA Ann Meet 2004

```
(6) Anon; ICS/IUGA Ann Meet 2004
(7) Anon; ICS/IUGA Ann Meet 2004
(8) Anon; ICS/IUGA Ann Meet 2004
(9) Anon; Rev Urol 2003, V5(1), P22
(10) Anon; Schwarz Pharma Annual Report 2003
(11) Breidenbach, A; 32nd Annu Meet Int Continence Soc, Abst 448 2002
(12) Cawello, W; Naunyn-Schmied Arch Pharmacol, Abst 428 2002, V365(Suppl 1)
(13) Chapple, C; XIXth Cong Eur Assoc Urol, Abst 508 2004
(14) Chapple, C; XIXth Cong Eur Assoc Urol, Abst 512 2004
(15) Eglen, R; Curr Opin Chem Biol 1999, V3, P426 CAPLUS
(16) Milsom, I; BJU International 2001, V87, P760 MEDLINE
(17) Prous Science Drug R & D Backgrounders; Urinary incontinence 2003
(18) Sachse, R; 32nd Annu Meet Int Continence Soc, Abst 440 2002
(19) Sachse, R; Eur Urol Suppl, Abst 111 2003, V2(1)
(20) Sachse, R; Naunyn-Schmied Arch Pharmacol, Abst 413 2002, V365(Suppl 1)
(21) Sachse, R; Naunyn-Schmied Arch Pharmacol, Abst 446 2003, V367(Suppl 1)
(22) Sachse, R; Proceedings of the International Continence Society 2003, V33,
    P377
(23) Sullivan, J; Eur Urol 1999, V36(Suppl 1), P89
(24) Wesnes, K; XIXth Cong Eur Assoc Urol, Abst 513 2004
L5
     ANSWER 6 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
     2004:872676 CAPLUS
AN
DN
     141:337790
ED
     Entered STN: 21 Oct 2004
     Transdermal administration of (R)-3,3-diphenylpropylamine monoesters
     Breitenbach, Armin; Meese, Claus; Wolff, Hans-Michael; Drews, Roland
IN
PA
     Schwarz Pharma Ag, Germany
SO
     PCT Int. Appl., 68 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     German
     ICM A61K009-70
IC
     ICS A61K031-403; C07C219-26
     63-6 (Pharmaceuticals)
     Section cross-reference(s): 1
FAN.CNT 1
                                            APPLICATION NO.
     PATENT NO.
                         KIND
                                DATE
                                                                   DATE
     -----
                         ----
                                _____
                                            ______
                                                                    _____
     WO 2004089346
                         A1
                                20041021
                                            WO 2004-EP3574
                                                                    20040403
PI
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
            CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
            GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
            LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
            NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
            TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
        RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
            BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,
            ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI,
            SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN,
            TD, TG
                                                                    20030408
     DE 10315878
                          Al
                                20041104
                                            DE 2003-10315878
                                                                    20040403
     AU 2004228927
                          A1
                                20041021
                                            AU 2004-228927
     CA 2505780
                         AA
                                20041021
                                            CA 2004-2505780
                                                                    20040403
                                                                    20040403
                         A1
                                20050518
                                            EP 2004-725614
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK,
     BR 2004006212
                         Α
                                20050816
                                            BR 2004-6212
                                                                    20040403
     CN 1767820
                                20060503
                                            CN 2004-80009176
                                                                    20040403
                         Α
     JP 2006522759
                         T2
                                20061005
                                            JP 2006-504992
                                                                    20040403
     ZA 2005002681
                                20051013
                                            ZA 2005-2681
                                                                    20050401
                         Α
     US 2006029673
                         A1
                                20060209
                                            US 2005-533683
                                                                    20050426
    NO 2005004644
                         Α
                                20051010
                                            NO 2005-4644
                                                                    20051010
PRAI DE 2003-10315878
                                20030408
                         Α
    WO 2004-EP3574
                         W
                                20040403
```

```
CLASS
                       PATENT FAMILY CLASSIFICATION CODES
 PATENT NO.
                 CLASS
                        _____
 _____
                 ----
 WO 2004089346
                 ICM
                        A61K009-70
                        A61K031-403; C07C219-26
                 ICS
                        A61K0009-70 [ICM,7]; A61K0031-403 [ICS,7]; C07C0219-26
                 IPCI
                        [ICS, 7]; C07C0219-00 [ICS, 7, C*]
                 IPCR
                        A61K0009-70 [I,C*]; A61K0009-70 [I,A]; A61K0031-403
                        [I,C*]; A61K0031-403 [I,A]
                        A61K009/70E; A61K031/403
                 ECLA
                        A61L0015-44 [ICM, 7]; A61L0015-16 [ICM, 7, C*]
 DE 10315878
                 IPCI
                        A61K0009-70 [I,C*]; A61K0009-70 [I,A]; A61K0031-403
                 IPCR
                        [I,C*]; A61K0031-403 [I,A]
                 ECLA
                        A61K009/70E; A61K031/403
                        A61K0009-70 [ICM,7]; A61K0031-403 [ICS,7]; C07C0219-26
 AU 2004228927
                IPCI
                        [ICS,7]; C07C0219-00 [ICS,7,C*]
                 IPCR
                        A61K0009-70 [I,C*]; A61K0009-70 [I,A]; A61K0031-403
                        [I,C*]; A61K0031-403 [I,A]
 CA 2505780
                 IPCI
                        A61K0009-70 [ICM,7]; C07C0219-26 [ICS,7]; C07C0219-00
                        [ICS,7,C*]; A61K0031-403 [ICS,7]
                        A61K0009-70 [I,C*]; A61K0009-70 [I,A]; A61K0031-403
                 IPCR
                        [I,C^*]; A61K0031-403 [I,A]
                 ECLA
                        A61K009/70E; A61K031/403
                        A61K0009-70 [ICM, 7]; A61K0031-403 [ICS, 7]; C07C0219-26
                 IPCI
 EP 1530461
                        [ICS,7]; C07C0219-00 [ICS,7,C*]
                        A61K0009-70 [I,C*]; A61K0009-70 [I,A]; A61K0031-403
                 IPCR
                        [I,C*]; A61K0031-403 [I,A]
                        A61K0009-70 [ICM,7]; A61K0031-403 [ICS,7]; C07C0219-26
                 IPCI
 BR 2004006212
                        [ICS,7]; C07C0219-00 [ICS,7,C*]
                        A61K0009-70 [I,C*]; A61K0009-70 [I,A]; A61K0031-403
                 IPCR
                        [I,C*]; A61K0031-403 [I,A]
                        A61K009/70E; A61K031/403
                 ECLA
                        A61K0009-70 [I,A]; A61K0031-403 [I,A]; C07C0219-26
 CN 1767820
                 IPCI
                        [I,A]; C07C0219-00 [I,C*]
                 ECLA
                        A61K009/70E; A61K031/403
                        A61K0031-222 [I,A]; A61K0031-21 [I,C*]; A61K0009-70
 JP 2006522759
                 IPCI
                        [I,A]; A61K0047-32 [I,A]; A61P0013-10 [I,A];
                        A61P0013-00 [I,A]
                        4C076/AA74; 4C076/BB31; 4C076/CC17; 4C076/EE08A;
                 FTERM
                        4C076/EE10A; 4C076/EE12A; 4C076/EE27A; 4C076/FF31;
                        4C076/FF68; 4C206/AA01; 4C206/AA02; 4C206/DB03;
                        4C206/DB04; 4C206/DB57; 4C206/MA01; 4C206/MA04;
                        4C206/MA52; 4C206/MA83; 4C206/NA11; 4C206/NA12;
                        4C206/ZA81
                        A61K [ICS, 7]; C07C [ICS, 7]
                 IPCI
 ZA 2005002681
                        A61K0009-70 [I,C*]; A61K0031-403 [I,C*]; A61K0009-70
                 IPCR
                        [I,A]; A61K0031-403 [I,A]
                 ECLA
                        A61K009/70E; A61K031/403
                        A61K0009-14 [I,A]
 US 2006029673
                 IPCI
                        424/486.000
                 NCL
                 ECLA
                        A61K009/70E; A61K031/403
                        A61K0009-70 [ICM,7]; A61K0031-403 [ICS,7]; C07C0219-26
 NO 2005004644
                 IPCI
                        [ICS,7]; C07C0219-00 [ICS,7,C*]
                        A61K0009-70 [I,C*]; A61K0009-70 [I,A]; A61K0031-403
                 IPCR
                        [I,C*]; A61K0031-403 [I,A]
                        A61K009/70E; A61K031/403
                 ECLA
OS
    MARPAT 141:337790
```

GI

Ozocerite

USES (Uses)

Polyoxyalkylenes, biological studies

IT

IT

The invention relates to a device for transdermally administering a compound AB of formula (I), wherein A represents hydrogen or deuterium, R represents a group selected among C1-6 alkyl, C3-10 cycloalkyl, or Ph, each of which can be substituted by C1-3 alkoxy, fluoride, chlorine, bromine, iodine, nitro, amino, hydroxy, oxo, mercapto, or deuterium, the C atom marked by * (asterisk) being provided in the R configuration. The invention is characterized in that the compound of general formula (I) is provided in a polymer matrix and is released at a dose of 0.5 to 20 mg per day through human skin. The invention further relates to the use of said compds. of formula (I) for producing transdermal medicaments. Thus a silicone-based transdermal system was prepared by the hot-melt process. 8.5 G of an adhesive mixture composed of BIO-PSA 7-4300 from Dow-Corning and 5 weight/weight% ozokerite or ceresin was heated to 150°C for 20 min until a homogeneous melt was formed. 1.5 G fesoterodine were added to the melt; the mixture was kept for addnl. 5 min at 150°C; followed by application onto a preheated foil. 5 Cm2 samples were used for dissoln. studies. transdermal diphenylpropylamine monoester Fesoterodineincontinence ST IT Isoprene-styrene rubber RL: PEP (Physical, engineering or chemical process); PYP (Physical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (block, triblock; transdermal administration of (R)-3,3diphenylpropylamine monoesters) IT Bladder, disease (incontinence; transdermal administration of (R)-3,3diphenylpropylamine monoesters) Urinary system, disease IT (nocturia; transdermal administration of (R)-3,3-diphenylpropylamine monoesters) IT Paraffin oils RL: PEP (Physical, engineering or chemical process); PYP (Physical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (ondina oil; transdermal administration of (R)-3,3-diphenylpropylamine monoesters) IT Dissolution Human Hydrophilicity

(transdermal administration of (R)-3,3-diphenylpropylamine monoesters)

(transdermal administration of (R)-3,3-diphenylpropylamine monoesters)

process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);

RL: PEP (Physical, engineering or chemical process); PYP (Physical

```
(transdermal administration of (R)-3,3-diphenylpropylamine monoesters)
     Drug delivery systems
TT
        (transdermal; transdermal administration of (R)-3,3-diphenylpropylamine
        monoesters)
     Urinary system, disease
IT
        (urinary frequency; transdermal administration of (R)-3,3-
        diphenylpropylamine monoesters)
                   700836-36-8D, block, triblock
ΙT
     700836-36-8
     RL: PEP (Physical, engineering or chemical process); PYP (Physical
     process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);
     USES (Uses)
        (isoprene-styrene rubber; transdermal administration of
        (R)-3,3-diphenylpropylamine monoesters)
     286930-02-7P, Fesoterodine
    RL: PEP (Physical, engineering or chemical process); PRP (Properties); PYP
     (Physical process); SPN (Synthetic preparation); THU (Therapeutic use);
     BIOL (Biological study); PREP (Preparation); PROC (Process); USES (Uses)
        (transdermal administration of (R)-3,3-diphenylpropylamine monoesters)
     1617-18-1, Ethylvinylacetate 198292-68-1, DuroTak 387-2287
IT
                                    504415-91-2, BIO-PSA 7-4300
     346577-82-0, Regalite R 1090
     RL: PEP (Physical, engineering or chemical process); PYP (Physical
     process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);
     USES (Uses)
        (transdermal administration of (R)-3,3-diphenylpropylamine monoesters)
     380636-50-0P
                   769950-53-0P
IT
     RL: PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use);
     BIOL (Biological study); PREP (Preparation); USES (Uses)
        (transdermal administration of (R)-3,3-diphenylpropylamine monoesters)
TT
     79-30-1, Isobutyric acid chloride
                                         110-17-8, Fumaric acid, reactions
     207679-81-0
     RL: RCT (Reactant); RACT (Reactant or reagent)
        (transdermal administration of (R)-3,3-diphenylpropylamine monoesters)
     5586-73-2D, 3,3-Diphenylpropylamine, monoesters of
                                                         9003-20-7, PVAc
IT
     9003-39-8, PVP
                     25322-68-3, PEO
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (transdermal administration of (R)-3,3-diphenylpropylamine monoesters)
RE.CNT · 6
              THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD
(1) Arth, C; US 6555129 B1 2003 CAPLUS
(2) Ebert, C; US 2002147236 A1 2002 CAPLUS
(3) Kanios, D; US 6638528 B1 2003 CAPLUS
(4) Sanol Arznei Schwarz Gmbh; EP 0957073 A 1999 CAPLUS
(5) Sanol Arznei Schwarz Gmbh; WO 0135957 A 2001
(6) Tsung-Min, H; US 2003157156 A1 2003 CAPLUS
    ANSWER 7 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
L5
     2004:761399 CAPLUS
AN
DN
     141:254396
     Entered STN: 19 Sep 2004
ED
     Fesoterodine a new effective and well-tolerated antimuscarinic
     for the treatment of urgency-frequency syndrome: results of a phase 2
     controlled study
CS
     Chapple C1, Royal Hallamshire Hospital, UK
     Neurourology and Urodynamics (2004), 23(5/6), 598-599
     CODEN: NEUREM; ISSN: 0733-2467
PB
     Wiley-Liss, Inc.
DT
     Journal
LΑ
     English
CC
     1-11 (Pharmacology)
     Fesoterodine as new effective and well-tolerated antimuscarinic
AB
     for the treatment of urgency-frequency syndrome is studied here.
ST
     antimuscarinic fesoterodine urgency frequency syndrome urinary
     incontinence
IT
     Human
```

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)

Muscarinic antagonists (antimuscarinic fesoterodine for treatment of urgency-frequency syndrome) IT Bladder, disease (incontinence; antimuscarinic fesoterodine for treatment of urgency-frequency syndrome) IT Disease, animal (urgency-frequency syndrome; antimuscarinic fesoterodine for treatment of urgency-frequency syndrome) IT 286930-02-7, Fesoterodine RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (antimuscarinic fesoterodine for treatment of urgency-frequency syndrome) ANSWER 8 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN L_5 2003:993805 CAPLUS ANDN 140:331551 Entered STN: 22 Dec 2003 ED Fesoterodine: Treatment of urinary incontinence muscarinic M3 TΙ antagonist ΑU Sorbera, L. A.; Castaner, J.; Lesson, P. A. CS Prous Science, Barcelona, 08080, Spain Drugs of the Future (2003), 28(7), 647-651 SO CODEN: DRFUD4; ISSN: 0377-8282 PB Prous Science DT Journal; General Review English LΑ CC1-0 (Pharmacology) A review. Urinary incontinence and overactive bladder are extremely AB common disorders affecting up to 12 and 20 million adults in the U.S., resp. Current pharmacotherapy includes peripherally acting compds. which modulate bladder smooth muscle contraction or centrally acting agents which modulate the neurol. control of urination. Anticholinergic agents inhibit bladder smooth muscle contraction through interference with acetylcholine action on muscarinic receptors on detrusor smooth muscle. However, the first anticholinergic agents were associated with a high rate of adverse events due to nonselectivity and targeting of several muscarinic subtypes and thus other organs. The search for novel, more bladder-selective antimuscarinic agents with better tolerability was initiated. Fesoterodine is a novel selective muscarinic M3 receptor antagonist that has shown potent antimuscarinic activity in vitro and in vivo and has been selected for further development as a treatment for urinary incontinence and overactive bladder. review fesoterodine urine incontinence muscarinic M3 antagonist ST IT Muscarinic antagonists (M3; fesoterodine treatment of urinary incontinence as muscarinic M3 antagonist) Bladder, disease IT (incontinence; fesoterodine treatment of urinary incontinence as muscarinic M3 antagonist) 286930-02-7, Fesoterodine IT RL: DMA (Drug mechanism of action); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (fesoterodine treatment of urinary incontinence as muscarinic M3 antagonist) THERE ARE 14 CITED REFERENCES AVAILABLE FOR THIS RECORD RE.CNT RE (1) Andersson, K; BJU Int 1999, V84, P923 CAPLUS

- (2) Andersson, K; Bailliere's Best Pract Res Clin Obstet Gynaecol 2000, V14, P291 MEDLINE
- (3) Anon; Schwarz's year-end results exceed expectations, DailyDrugNews com 2003
- (4) Breidenbach, A; 32nd Annu Meet Int Continence Soc Abst 448 2002
- (5) Eglen, R; Curr Opin Chem Biol 1999, V3, P426 CAPLUS

```
(6) Meese, C; WO 0135957
(7) Meese, C; DE 19955190 CAPLUS
(8) Meese, C; JP 2003514018 CAPLUS
(9) Prous Science Drug R&d; Backgrounders: Urinary incontinence (online
   publication) 2003
(10) Sachse, R; 32nd Annu Meet Int Continence Soc Abst 440 2002
(11) Sachse, R; Eur Urol Suppl Abst 111 2003, V2(1)
(12) Sachse, R; Naunyn-Schmied Arch Pharmacol Abst 413 2002, V365(Suppl 1)
(13) Sachse, R; Naunyn-Schmied Arch Pharmacol Abst 446 2003, V367(Suppl 1)
(14) Sullivan, J; Eur Urol 1999, V36(Suppl 1), P89
    ANSWER 9 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN
L5
ΑN
    2003:950829 CAPLUS
DN
    140:13084
    Entered STN: 07 Dec 2003
ED
    Combination of selected opioids with other active substances for use in
TI
    the therapy of urinary incontinence
    Christoph, Thomas
IN
    Grunenthal G.m.b.H., Germany
PA
    PCT Int. Appl., 126 pp.
SO
    CODEN: PIXXD2
DT
    Patent
LA
    German
IC
    ICM A61K031-135
    ICS A61K031-137; A61K031-485
    1-12 (Pharmacology)
CC
    Section cross-reference(s): 63
FAN.CNT 1
                                         APPLICATION NO. DATE
    PATENT NO.
                       KIND DATE
    WO 2003099268 A1 20031204 WO 2003-EP5529 20030527
PΙ
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CO, CR, CU, CZ, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM,
            HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS,
            LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL,
            PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA,
            UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
            KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES,
            FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                               20031211 DE 2002-10224107 20020529
                        A1
    DE 10224107
                                                                20030527
                                        AU 2003-240717
EP 2003-730120
    AU 2003240717
                         A1
                               20031212
                         A1
                               20050223
                                                                 20030527
    EP 1507520
           AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
                                         US 2004-998164 20041129
                   A1
                               20050623
    US 2005137194
                                           US 2005-545901
                                                                 20050817
                         A1
                               20060803
    US 2006168942
PRAI DE 2002-10224107
                        Α
                               20020529
    WO 2003-EP5529
                         W
                               20030527
CLASS
                CLASS PATENT FAMILY CLASSIFICATION CODES
PATENT NO.
                       _____
                ----
 ______
WO 2003099268
                ICM
                       A61K031-135
                       A61K031-137; A61K031-485
                ICS
                IPCI
                       A61K0031-135 [ICM,7]; A61K0031-137 [ICS,7];
                       A61K0031-485 [ICS,7]
                       A61K0031-135 [I,C*]; A61K0031-135 [I,A]; A61K0031-137
                IPCR
                       [I,C*]; A61K0031-137 [I,A]; A61K0031-485 [I,C*];
                       A61K0031-485 [I,A]
                ECLA
                       A61K031/135; A61K031/137; A61K031/485
                       A61K0031-485 [ICM, 7]
DE 10224107
                IPCI
                       A61K0031-135 [I,C*]; A61K0031-135 [I,A]; A61K0031-137
                IPCR
                       [I,C*]; A61K0031-137 [I,A]; A61K0031-485 [I,C*];
                       A61K0031-485 [I,A]
```

```
A61K031/135; A61K031/137; A61K031/485
                 ECLA
                 IPCI
                        A61K0031-135 [ICM,7]; A61K0031-137 [ICS,7];
 AU 2003240717
                        A61K0031-485 [ICS,7]
                        A61K0031-135 [I,C*]; A61K0031-135 [I,A]; A61K0031-137
                 IPCR
                        [I,C*]; A61K0031-137 [I,A]; A61K0031-485 [I,C*];
                        A61K0031-485 [I,A]
 EP 1507520
                 IPCI
                        A61K0031-135 [ICM, 7]; A61K0031-137 [ICS, 7];
                        A61K0031-485 [ICS, 7]
                 IPCR
                        A61K0031-135 [I,C*]; A61K0031-135 [I,A]; A61K0031-137
                        [I,C*]; A61K0031-137 [I,A]; A61K0031-485 [I,C*];
                        A61K0031-485 [I,A]
 US 2005137194
                 IPCI
                        A61K0031-5377 [ICM,7]; A61K0031-5375 [ICM,7,C*];
                        A61K0031-485 [ICS, 7]
                 IPCR
                        A61K0031-485 [I,C*]; A61K0031-485 [I,A]; A61K0031-5375
                        [I,C*]; A61K0031-5377 [I,A]
                        514/235.200; 514/282.000
                 NCL
 US 2006168942
                        F01N0009-00 [I,A]
                 IPCI
                 NCL
                        060/276.000; 060/285.000
                 ECLA
                        A61K031/135; A61K031/137; A61K031/485
    MARPAT 140:13084
os
     The invention discloses the use of a combination of opioids (e.g.
AB
     tramadol) with other active substances for producing a drug for the
     treatment of urinary urgency or urinary incontinence. The invention also
     relates to corresponding medicaments and to a method for treating urinary
     urgency or urinary incontinence.
     incontinence urinary treatment opioid drug combination; urinary urge
ST
     treatment opioid drug combination; tramadol drug combination urinary
     incontinence urge
TT
     Bladder, disease
        (incontinence; opioid combination with other active substances for
        treatment of urinary incontinence)
     Drug delivery systems
IT
        (injections; opioid combination with other active substances for
        treatment of urinary incontinence)
IT
    Drug delivery systems
        (opioid combination with other active substances for treatment of
        urinary incontinence)
    Bladder
ΙT
        (urinary urge; opioid combination with other active substances for
        treatment of urinary incontinence)
    57-27-2, * Morphin, biological studies
                                              57-42-1, Pethidine
IT
                 76-42-6, Oxycodone
                                      76-57-3, Codeine
                                                         76-58-4,
    Nalorphine
                                           125-28-0, Dihydrocodeine
                    77-07-6, Levorphanol
    Ethylmorphine
     125-29-1, Hydrocodone 125-58-6, Levomethadone 302-41-0, Piritramide
     357-56-2, Dextromoramide 359-83-1, Pentazocine
                                                        437-38-7, Fentanyl
     466-99-9, Hydromorphone 469-62-5, Dextropropoxyphene
                                                              469-79-4,
                   561-27-3, Diacetylmorphine
                                                915-30-0, Diphenoxylate
    Ketobemidone
                           1477-40-3, Levomethadyl Acetate
                                                            14521-96-1,
    1199-99-1D, derivs.
                                          21363-18-8, Viminol
                                                                27203-92-5,
                20594-83-6, Nalbuphine
    Etorphine
                42408-82-2, Butorphanol
                                          51931-66-9, Tilidine
                                                                52485-79-7,
    Tramadol
                     53648-55-8, Dezocine
                                            54340-58-8, Meptazinol
                                                                     56030-54-7
    Buprenorphine
     71195-58-9, Alfentanyl
                             80456-81-1, O-Demethyltramadol
                                                               132875-61-7,
    Remifentanyl
                   138853-73-3
    RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (Combination of selected opioids with other active substances for use
        in the therapy of urinary incontinence)
IT
    186033-14-7, NS 8
    RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (NS 8; opioid combination with other active substances for treatment of
       urinary incontinence)
IT
    52-28-8, Codeine phosphate
                                  57444-62-9, Resiniferatoxin
                                                                92725-18-3D,
              93413-69-5, Venlafaxine 142155-43-9, Cizolirtine
    158836-71-6, Nitro-Flurbiprofen 174636-32-9, Talnetant
                                                                175590-75-7
```

175590-76-8 175590-77-9 175590-78-0 175590-89-3 175590-90-6 175590-91-7 175591-02-3 175591-01-2 175590-92-8 175591-04-5 175591-05-6 175591-09-0 175591-11-4 175591-06-7 175591-12-5 175591-25-0 175591-23-8 175591-24-9 175774-12-6 175774-14-8 175774-16-0 175774-18-2 187219-61-0 187219-93-8 187219-95-0 187219-97-2 187219-99-4 187220-01-5 187220-05-9 187220-25-3 217185-75-6, TAK-637 219311-44-1 220382-87-6, Rec 187220-29-7 15/3079 242478-37-1, Solifenacin 286930-02-7, Fesoterodine 433265-59-9 433265-42-0 433265-54-4 433265-65-7 433265-73-7 433686-04-5 433686-06-7 433686-05-6 433686-07-8 433936-14-2 433936-20-0 433936-23-3 433936-24-4 502616-18-4 502616-19-5 502616-23-1 502616-20-8 502616-22-0 630046-59-2 630395-07-2, SL 251039 630395-09-4, DRP 001

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(opioid combination with other active substances for treatment of urinary incontinence)

RE.CNT 9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD RE

- (1) Durand, A; PRESSE MEDICALE 2000, V29(16), P917
- (2) Gruenenthal Gmbh; DE 19947747 A 2001 CAPLUS
- (3) Kroner, B; JOURNAL OF GERIATRIC DRUG THERAPY 1992, V7(1), P23
- (4) Malinovsky, J; ANESTHESIA AND ANALGESIA 1998, V87(2), P456 CAPLUS
- (5) McNutt, R; US 5658908 A 1997 CAPLUS
- (6) Novosis Pharma Ag; EP 1072260 A 2001 CAPLUS
- (7) Palmer, K; GASTROENTEROLOGY 1980, V79(6), P1272 MEDLINE
- (8) Pandita, R; NEUROUROLOGY AND URODYNAMICS, 31st Annual Meeting of the International Continence Society 2001, V20(4), P439
- (9) Ripple, M; AMERICAN JOURNAL OF FORENSIC MEDICINE AND PATHOLOGY 2000, V21(4), P370 MEDLINE

=> FIL MARPAT

| COST IN U.S. DOLLARS | SINCE FILE | TOTAL |
|--|----------------|------------------|
| • | ENTRY | SESSION |
| FULL ESTIMATED COST | 32.45 | 61.58 |
| DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) | SINCE FILE | TOTAL SESSION |
| CA SUBSCRIBER DRICE | ENTRY -6.75 | |

FILE 'MARPAT' ENTERED AT 14:42:35 ON 02 NOV 2006
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
COPYRIGHT (C) 2006 American Chemical Society (ACS)

FILE CONTENT: 1961-PRESENT VOL 145 ISS 18 (20061027/ED)

SOME MARPAT RECORDS ARE DERIVED FROM INPI DATA FOR 1961-1987

MOST RECENT CITATIONS FOR PATENTS FROM MAJOR ISSUING AGENCIES (COVERAGE TO THESE DATES IS NOT COMPLETE):

```
7108861 19 SEP 2006
DE 102006006123 07 SEP 2006
EΡ
        1700848 13 SEP 2006
JΡ
     2006242783 14 SEP 2006
     2006095864 14 SEP 2006
WO
GB
        2423518 30 AUG 2006
FR
        2882520 01 SEP 2006
RU
        2283369 10 SEP 2006
CA
        2547866 22 AUG 2006
```

Expanded G-group definition display now available.

=> SET NOTICE DISPLAY 1

NOTICE SET TO 1 U.S. DOLLAR FOR DISPLAY COMMAND SET COMMAND COMPLETED

treatment of urinary incontinence)

Drug delivery systems

urinary incontinence)

IT

=> D ACC 140:13084 ALL

THE ESTIMATED COST FOR THIS REQUEST IS 4.76 U.S. DOLLARS DO YOU WANT TO CONTINUE WITH THIS REQUEST? (Y)/N:y

```
ANSWER 1 MARPAT COPYRIGHT 2006 ACS on STN
    140:13084 MARPAT
AN
    Combination of selected opioids with other active substances for use in
TT
    the therapy of urinary incontinence
    Christoph, Thomas
IN
    Grunenthal G.m.b.H., Germany
PA
    PCT Int. Appl., 126 pp.
SO
    CODEN: PIXXD2
    Patent
DT
LA
    German
    ICM A61K031-135
IC
    ICS A61K031-137; A61K031-485
CC
    1-12 (Pharmacology)
    Section cross-reference(s): 63
FAN.CNT 1
                                          APPLICATION NO. DATE
    PATENT NO.
                     KIND DATE
                     ----
                                          -----
     _____
                                        WO 2003-EP5529 20030527
PΤ
    WO 2003099268
                    A1 20031204
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CO, CR, CU, CZ, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM,
            HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS,
            LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL,
            PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA,
            UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
            KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES,
            FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR,
            BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
    DE 10224107
                     A1
                           20031211
                                        DE 2002-10224107 20020529
    AU 2003240717
                                         AU 2003-240717
                                                           20030527
                      A1
                           20031212
                                         EP 2003-730120
                                                           20030527
    EP 1507520
                      Α1
                          20050223
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
    US 2005137194
                                        US 2004-998164
                                                           20041129
                     A1
                           20050623
                                         US 2005-545901
                                                           20050817
    US 2006168942
                      A1
                           20060803
PRAI DE 2002-10224107 20020529
                     20030527
    WO 2003-EP5529
    The invention discloses the use of a combination of opioids (e.g.
AΒ
    tramadol) with other active substances for producing a drug for the
    treatment of urinary urgency or urinary incontinence. The invention also
    relates to corresponding medicaments and to a method for treating urinary
    urgency or urinary incontinence.
ST
    incontinence urinary treatment opioid drug combination; urinary urge
    treatment opioid drug combination; tramadol drug combination urinary
    incontinence urge
IT
    Bladder, disease
        (incontinence; opioid combination with other active substances for
        treatment of urinary incontinence)
IT
    Drug delivery systems
        (injections; opioid combination with other active substances for
```

(opioid combination with other active substances for treatment of

```
IT
    Bladder
        (urinary urge; opioid combination with other active substances for
        treatment of urinary incontinence)
    57-27-2, * Morphin, biological studies 57-42-1, Pethidine
IT
                 76-42-6, Oxycodone
                                      76-57-3, Codeine
                                                          76-58-4,
    Nalorphine
    Ethylmorphine
                     77-07-6, Levorphanol
                                           125-28-0, Dihydrocodeine
                             125-58-6, Levomethadone
                                                       302-41-0, Piritramide
    125-29-1, Hydrocodone
     357-56-2, Dextromoramide
                               359-83-1, Pentazocine
                                                        437-38-7, Fentanyl
                               469-62-5, Dextropropoxyphene
                                                              469-79-4,
     466-99-9, Hydromorphone
                   561-27-3, Diacetylmorphine
                                                915-30-0, Diphenoxylate
    Ketobemidone
     1199-99-1D, derivs.
                           1477-40-3, Levomethadyl Acetate
                                                             14521-96-1,
                20594-83-6, Nalbuphine
                                         21363-18-8, Viminol
    Etorphine
                                                                27203-92-5,
               42408-82-2, Butorphanol
                                          51931-66-9, Tilidine
                                                                 52485-79-7,
    Tramadol
    Buprenorphine
                     53648-55-8, Dezocine 54340-58-8, Meptazinol
                                                                     56030-54-7
                              80456-81-1, O-Demethyltramadol
                                                               132875-61-7,
     71195-58-9, Alfentanyl
                   138853-73-3
    Remifentanyl
    RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (Combination of selected opioids with other active substances for use
        in the therapy of urinary incontinence)
    186033-14-7, NS 8
IT
    RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (NS 8; opioid combination with other active substances for treatment of
       urinary incontinence)
    52-28-8, Codeine phosphate
                                  57444-62-9, Resiniferatoxin
                                                                92725-18-3D,
IT
              93413-69-5, Venlafaxine 142155-43-9, Cizolirtine
                                                                175590-75-7
     158836-71-6, Nitro-Flurbiprofen
                                     174636-32-9, Talnetant
                                               175590-89-3
     175590-76-8
                  175590-77-9
                                 175590-78-0
                                                             175590-90-6
                                                             175591-04-5
     175590-91-7
                  175590-92-8
                                 175591-01-2
                                               175591-02-3
                                                             175591-12-5
                                 175591-09-0
                                               175591-11-4
    175591-05-6
                  175591-06-7
                                               175774-12-6
                                                             175774-14-8
                                 175591-25-0
     175591-23-8
                  175591-24-9
                                               187219-93-8
                                                             187219-95-0
                                 187219-61-0
     175774-16-0
                 175774-18-2
                                 187220-01-5
                                               187220-05-9
                                                             187220-25-3
    187219-97-2
                  187219-99-4
                                                        220382-87-6, Rec
    187220-29-7
                  217185-75-6, TAK-637
                                          219311-44-1
                                          286930-02-7, Fesoterodine
             242478-37-1, Solifenacin
    15/3079
                                               433265-65-7
                                                             433265-73-7
                                 433265-59-9
     433265-42-0
                  433265-54-4
                                                             433936-14-2
                                               433686-07-8
    433686-04-5
                   433686-05-6
                                 433686-06-7
                                 433936-24-4
                                               502616-18-4
                                                             502616-19-5
     433936-20-0
                   433936-23-3
     502616-20-8
                   502616-22-0
                                 502616-23-1
                                               630046-59-2
                                                             630395-07-2, SL
    251039
              630395-09-4, DRP 001
    RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (opioid combination with other active substances for treatment of
       urinary incontinence)
              THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
(1) Durand, A; PRESSE MEDICALE 2000, V29(16), P917
(2) Gruenenthal Gmbh; DE 19947747 A 2001 CAPLUS
(3) Kroner, B; JOURNAL OF GERIATRIC DRUG THERAPY 1992, V7(1), P23
(4) Malinovsky, J; ANESTHESIA AND ANALGESIA 1998, V87(2), P456 CAPLUS
(5) McNutt, R; US 5658908 A 1997 CAPLUS
(6) Novosis Pharma Ag; EP 1072260 A 2001 CAPLUS
(7) Palmer, K; GASTROENTEROLOGY 1980, V79(6), P1272 MEDLINE
(8) Pandita, R; NEUROUROLOGY AND URODYNAMICS, 31st Annual Meeting of the
   International Continence Society 2001, V20(4), P439
(9) Ripple, M; AMERICAN JOURNAL OF FORENSIC MEDICINE AND PATHOLOGY 2000,
```

V21(4), P370 MEDLINE

G1 = OH / F / Cl / H / 22

- G2 = carbon chain <containing 1-3 C> (opt. substd.) /
- (Specifically claimed: Me)
- G3 = carbon chain <containing 1-4 C> (opt. substd.) / (Specifically claimed: Me / Et / Bu-n / Bu-t)

- G5 = H / carbon chain <containing 1-4 C> (opt. substd.) / (Specifically claimed: Me / Et / Pr-i / Bu-t)
- G6 = 2-136 3-134 4-135 6-12 / 46-136 47-134 48-135 49-12 / 55-136 59-134 58-135 60-12

G7 = H / F / Cl / Br / I / 29 / OH / SH / 33 / OCF3 / NH2 / 35 / SO2Me / SO2CF3 / CN / 91 / NO2 / CONH2 / 41 / carbon chain <containing 1-6 C> (opt. substd.) / Ph (opt. substd.)

- G8 = O / S
- G9 = carbon chain <containing 1-6 C> (opt. substd.) /
 pyridyl / thienyl / thiazolyl / Ph / CH2Ph / CH2CH2Ph / 101 /
 142 / 105 / 146 / 110 / 116 / 124 /
 (Specifically claimed: Me)

G10 = NH / 37

Gl1 = carbon chain <containing 1-6 C> (opt. substd.) /
Ph (opt. substd.) / CH2Ph (opt. substd.) /
CH2CH2Ph (opt. substd.)

G12 = H / F

G13 = OCH2O / OCH2CH2O / 66-45 68-44 / 69-45 71-44 / CH2CH2CH2CH2 / 74-45 77-44

G14 = H / Me G15 = OCH2O / OCH2CH2O / 78-56 80-57 / 82-56 84-57 / CH2CH2CH2CH2 / 86-56 89-57

G16 = carbon chain <containing 1-4 C> (opt. substd.)

G17 = phenylene

G18 = alkyl <containing 1-3 C>

G19 = H / carbon chain <containing 1-6 C> (opt. substd.) / Ph (opt. substd.) / CH2Ph (opt. substd.) /

CH2CH2Ph (opt. substd.) = carbon chain <containing 1-3 C> (opt. substd.)

G20 = carbon chain <contain G21 = m-C6H4 / p-C6H4

G22 = 132 / morpholino

G23 = carbon chain <containing 1-4 C> (opt. substd.)

G24 = NH2 / 137

HN-G11

G25 = carbon chain <containing 1-5 C> (opt. substd.)

G26 = 148 / 151

$$G1$$
 C CH_2 Me $G1$ CH_2 Me $G3$ 148 $G4$ N $G27$ Me Me

G27 = alkyl <containing 1-4 C> / CH2Ph / CF3 / OH / OCH2Ph / alkoxy <containing 1-4 C> / Cl / F / H /

(Specifically claimed: Me)

Patent location:

claim 1

Note:

and/or physiologically acceptable salts

Stereochemistry: and enantiomers, diastereomers and mixtures

=> SET NOTICE LOGIN DISPLAY

NOTICE SET TO OFF FOR DISPLAY COMMAND SET COMMAND COMPLETED

=> FIL STNGUIDE

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION FULL ESTIMATED COST 8.14 69.72

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

SINCE FILE TOTAL
ENTRY SESSION

CA SUBSCRIBER PRICE -0.71 -7.46

FILE 'STNGUIDE' ENTERED AT 14:43:44 ON 02 NOV 2006
USE IS SUBJECT TO THE TERMS OF YOUR CUSTOMER AGREEMENT
COPYRIGHT (C) 2006 AMERICAN CHEMICAL SOCIETY, JAPAN SCIENCE
AND TECHNOLOGY CORPORATION, AND FACHINFORMATIONSZENTRUM KARLSRUHE

FILE CONTAINS CURRENT INFORMATION. LAST RELOADED: Oct 27, 2006 (20061027/UP).

=>

---Logging off of STN---

Executing the logoff script...

=> LOG Y

---Logging off of STN---

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION FULL ESTIMATED COST 0.30 70.02

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

SINCE FILE

E FILE TOTAL ENTRY SESSION

CA SUBSCRIBER PRICE

0.00

-7.46

STN INTERNATIONAL LOGOFF AT 14:46:58 ON 02 NOV 2006

Connection closed by remote host END

Unable to generate the STN prompt. Exiting the script...